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L1            1 US20020065296/PN OR US99-115878/AP,PRN

     FILE 'REGISTRY' ENTERED AT 09:20:28 ON 06 JUN 2005

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L2              TRA L1 1- RN :      157 TERMS

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L4            1 US20020065296/PN OR US99-115878/AP,PRN

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L1   ANSWER 1 OF 1  HCAPLUS  COPYRIGHT 2005 ACS on STN
AN   2002:409267  HCAPLUS
DN   137:6098
ED   Entered STN:  31 May 2002
TI   Heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors
IN   Dumas, Jacques; Riedl, Bernd; Khire, Uday; Sibley, Robert N.;
     Hatoum-Mokdad, Holia; Monahan, Mary-katherine; Gunn, David E.; Lowinger,
     Timotthy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.
PA   Bayer Corporation, USA
SO   U.S. Pat. Appl. Publ., 39 pp., Cont.-in-part of U. S. Ser. No. 778,039.
     CODEN: USXXCO
DT   Patent
LA   English
IC   ICM  A61K031-506
     ICS  A61K031-501; A61K031-497; A61K031-4725; A61K031-4709
INCL 514310000
CC   27-17 (Heterocyclic Compounds (One Hetero Atom))
     Section cross-reference(s): 1
FAN.CNT 5
     PATENT NO.          KIND   DATE        APPLICATION NO.        DATE
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PI   US 2002065296        A1    20020530    US 2001-838286         20010420 <--
     US 2003139605        A1    20030724    US 2002-71248          20020211
     CA 2443952           AA    20021031    CA 2002-2443952        20020417
     WO 2002085859        A1    20021031    WO 2002-US12064        20020417
        W:  AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL,
            PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG,
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            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     EP 1379507           A1    20040114    EP 2002-725709         20020417
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     JP 2004537511        T2    20041216    JP 2002-583386         20020417
PRAI US 1999-115878P      P     19990113
     US 1999-257265       B1    19990225
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     US 2001-778039       A2    20010207
     US 1999-115877P      P     19990113
     US 1999-257266       B2    19990225
     US 1999-425228       B1    19991022
     US 2001-838286       A     20010420
     US 2001-948915       A1    20010910
     WO 2002-US12064      W     20020417
CLASS
 PATENT NO.       CLASS  PATENT FAMILY CLASSIFICATION CODES
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 US 2002065296    ICM    A61K031-506
                  ICS    A61K031-501; A61K031-497; A61K031-4725; A61K031-4709
                  INCL   514310000
 US 2002065296    NCL    514/310.000; 514/313.000; 514/336.000; 514/337.000;
                         514/252.030; 514/252.040; 514/255.050; 514/256.000
                  ECLA   A61K031/17; A61K031/18; A61K031/24; A61K031/341;
                         A61K031/40+A; A61K031/4035; A61K031/44; A61K031/44+A;
                         A61K031/4439; A61K031/4453; A61K031/47; A61K031/4709;
                         A61K031/4725; A61K031/495+A; A61K031/496; A61K031/5375;
                         A61K031/5377; C07D213/75D3; C07D215/38C; C07D217/22;
                         C07D401/12+215+213                                <--
 US 2003139605    NCL    546/291.000
                  ECLA   A61K031/17; C07C311/29; C07C317/22; C07D209/48D5C1;
                         C07D213/75D3; C07D213/81E; C07D295/12A1;
                         C07D295/12B1D4; C07D295/18B2D; A61K031/18; A61K031/24;
                         A61K031/341; A61K031/40+A; A61K031/4035; A61K031/44+A;
                         A61K031/4439; A61K031/4453; A61K031/495+A; A61K031/496;
                         A61K031/5375; A61K031/5377; C07C275/28; C07C275/30;
                         C07C275/32; C07C275/36; C07C275/40
 WO 2002085859    ECLA   A61K031/44; A61K031/47; A61K031/4709; A61K031/4725;
                         C07D213/75D3; C07D215/38C; C07D217/22;
                         C07D401/12+215+213; C07D401/12+217+213
 JP 2004537511    FTERM  4C031/JA09; 4C034/AL05; 4C055/AA01; 4C055/BA01;
                         4C055/BA02; 4C055/BA53; 4C055/BB17; 4C055/CA01;
                         4C055/DA06; 4C055/DA28; 4C055/DA42; 4C055/DA47;
                         4C055/DB10; 4C055/DB17; 4C055/EA01; 4C063/AA01;
                         4C063/BB07; 4C063/BB09; 4C063/CC14; 4C063/CC15;
                         4C063/DD07; 4C063/DD12; 4C063/EE01; 4C086/AA01;
                         4C086/AA03; 4C086/BC17; 4C086/BC28; 4C086/BC30;
                         4C086/BC50; 4C086/GA07; 4C086/GA08; 4C086/GA12;
                         4C086/MA01; 4C086/MA04; 4C086/NA14; 4C086/ZA01;
                         4C086/ZA02; 4C086/ZA36; 4C086/ZA45; 4C086/ZA54;
                         4C086/ZA59; 4C086/ZA67; 4C086/ZA68; 4C086/ZA75;
                         4C086/ZA86; 4C086/ZA89; 4C086/ZA94; 4C086/ZA96;
                         4C086/ZA97; 4C086/ZB02; 4C086/ZB05; 4C086/ZB11;
                         4C086/ZB13; 4C086/ZB15; 4C086/ZB26; 4C086/ZB33;
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4C086/ZB35; 4C086/ZC21; 4C086/ZC35

OS MARPAT 137:6098

AB This invention relates to the use of a group of heteroaryl ureas (I; for example, N-(2-methoxy-3-quinolyl)-N'-[4-[3-(N-methylcarbamoyl)phenoxy]phenyl]urea) containing N in treating p38 mediated diseases, and pharmaceutical compns. for use in such therapy. I is A-NHC(O)NH-B or a pharmaceutically acceptable salt thereof, wherein A is a substituted or unsubstituted pyridyl, quinolinyl or isoquinolinyl group, B is a substituted or unsubstituted, up to tricyclic aryl or heteroaryl moiety of up to 50 C atoms with a cyclic structure bound directly to N, containing at least 5 cyclic members with 0-4 members of groups consisting of N, O and S. Information about the substituents for A and B are given in the claims. Although the methods of preparation are not claimed, 37 example prepns. are included as well as examples of preparation of intermediates. No pharmacol. data is included.

ST nitrogen heteroaryl urea prepn p38 kinase inhibitor; pyridyl urea prepn p38 kinase inhibitor; quinolyl urea prepn p38 kinase inhibitor; isoquinolyl urea prepn p38 kinase inhibitor

IT Infection
(Chagas' disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Inflammation
(Crohn's disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Intestine, disease
(Crohn's; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Disease, animal
(Jarisch-Herxheimer reaction; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Malaria
(Plasmodium falciparum malaria and cerebral malaria; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treating)

IT Antimalarials
(Plasmodium falciparum malaria and cerebral malaria; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Toxins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(Shiga-like toxin, effects of toxins from Escherichia coli infection; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Respiratory distress syndrome
(adult; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Hepatitis
(alc.; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Transplant rejection
(allotransplant; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Lung
(alveolus, injury; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Antiarteriosclerotics
(antiatherosclerotics; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Aneurysm
(aortic; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Meningitis
(bacterial; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Necrosis
(bowel; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38

kinase inhibitors for treatment of)
IT Bronchi, disease
Inflammation
(bronchitis, obliterative; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Injury
(cerebral; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Pneumoconiosis
(coal worker's; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Eye, disease
(cornea, ulcer; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Ulcer
(corneal; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Radiation
(damage, injury/toxicity following administration of monoclonal antibodies; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Cartilage, disease
(degeneration; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Multiple sclerosis
(demyelation and oligiodendrocyte loss in; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Liver, disease
(during acute inflammation; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Toxins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(enterotoxin A, effects of toxins from Staphylococcus infection; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Skin, disease
(epidermolysis bullosa, dystrophobic; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Liver, disease
(failure; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Lung, disease
(fibrosis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Nervous system agents
(for demyelating disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)
IT Wound healing
(impaired wound healing in infection; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Helicobacter pylori
(infection during peptic ulcer disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Borrelia burgdorferi
Cytomegalovirus
Human immunodeficiency virus
Influenza virus
Theiler's murine encephalomyelitis virus
Treponema pallidum
(infections from; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Brain, disease
Reperfusion
(injury; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Leukemia

(lymphocytic, inhibitors; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Neoplasm
(metastasis, inhibitors; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Heterocyclic compounds
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(nitrogen, heteroaryl ureas; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

IT Bone, disease
(osteopenia, mediated by MMP activity; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Inflammation
Pancreas, disease
(pancreatitis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Ulcer
(peptic, Helicobacter pylori infection during; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Osteoporosis
(postmenopausal; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treating)

IT Human
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

IT Allergy
Alzheimer's disease
Arthritis
Asthma
Diabetes mellitus
Rheumatoid arthritis
Tuberculosis
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treating)

IT Encephalitis
Myelodysplastic syndromes
Periodontium, disease
Psoriasis
Rheumatic fever
Silicosis
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Allergy inhibitors
Anti-Alzheimer's agents
Anti-infective agents
Anti-inflammatory agents
Antiarthritics
Antiasthmatics
Antibacterial agents
Anticoagulants
Antidiabetic agents
Antirheumatic agents
Antitumor agents
Cardiovascular agents
Contraceptives
Tuberculostatics
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Biliary tract, disease
(primary biliary cirrhosis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Proteins
RL: BSU (Biological study, unclassified); BIOL (Biological study)

(proteinuria; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Fibrosis
Sarcoidosis
(pulmonary; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Injury
(reperfusion; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Bone
(resorption; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Lung, disease
(sarcoidosis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Shock (circulatory collapse)
(septic; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Inflammation
(systemic inflammatory response syndrome; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Lupus erythematosus
(systemic; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Disease, animal
(temporomandibular joint; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Joint, anatomical
(temporomandibular, disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Osteoporosis
(therapeutic agents, postmenopausal; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Shock (circulatory collapse)
(toxic shock syndrome; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Digestive tract, disease
(ulcer, peptic, Helicobacter pylori infection during; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Inflammation
Intestine, disease
(ulcerative colitis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT 165245-96-5, p38 Kinase
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(inhibitors; preparation of heteroaryl ureas containing nitrogen hetero-atoms as)

IT 673-09-6P, 4-(4-Pyridylthio)aniline 6337-24-2P, 1-Methoxy-4-(4-nitrophenoxy)benzene 13472-85-0P, 5-Bromo-2-methoxypyridine 18994-90-6P, 4-(1-Imidazolylmethyl)-1-nitrobenzene 27237-21-4P, 4-(3-Carboxyphenoxy)-1-nitrobenzene 27692-74-6P, 4-(4-Pyridinylmethyl)aniline 28232-34-0P, 5-Nitro-2-(4-methylphenoxy)pyridine 28232-52-2P, 3-(3-Pyridinyloxy)-1-nitrobenzene 29264-35-5P, 4-(3-Carboxy-4-hydroxyphenoxy)-1-nitrobenzene 31465-36-8P, 4-(4-Methoxyphenoxy)aniline 32361-76-5P, 3-(4-Nitrobenzyl)pyridine 36089-89-1P, 4-(4-Methylsulfonylphenoxy)-1-nitrobenzene 50727-06-5P, 5-Hydroxyisoindoline-1,3-dione 51727-15-2P, 4-Chloropyridine-2-carbonyl chloride hydrochloride 51834-97-0P, 5-Hydroxy-2-methoxypyridine 56643-85-7P, 4-(1-Imidazolylmethyl)aniline 62248-47-9P, 4-[(4-Butoxyphenyl)thio]-1-nitrobenzene 62248-51-5P, 4-(4-Butoxyphenyl)thioaniline 64064-63-7P, 4-(6-Methyl-3-pyridinyloxy)-1-nitrobenzene 70991-08-1P, 4-(2-Pyridinylthio)aniline 71708-64-0P, 4-[3-(N-Methylcarbamoyl)phenoxy]-1-nitrobenzene 85666-15-5P, 4-[(3-Pyridinyl)methyl]aniline 92575-23-0P, 3-(4-Pyridinylthio)aniline 99586-65-9P, 4-Chloro-2-pyridinecarboxamide 102877-78-1P 116289-71-5P,

3-(3-Pyridinyloxy)aniline 135680-03-4P, 4-(4-tert-Butoxycarbonylaminobenzyl)aniline 176977-85-8P, Methyl 4-chloropyridine-2-carboxylate hydrochloride 178809-75-1P, 4-[Hydroxy(4-pyridyl)methyl]-1-nitrobenzene 220000-87-3P 228401-26-1P, 3-(Trifluoromethyl)-4-(4-pyridinylthio)nitrobenzene 228401-27-2P, 3-(Trifluoromethyl)-4-(4-pyridinylthio)aniline 228401-28-3P, 4-[(4-Phenyl-2-thiazolyl)thio]-1-nitrobenzene 228401-29-4P, 4-[(4-Phenyl-2-thiazolyl)thio]aniline 228401-31-8P, 4-(6-Methyl-3-pyridinyloxy)aniline 228401-32-9P, 4-(3,4-Dimethoxyphenoxy)-1-nitrobenzene 228401-33-0P, 4-(3,4-Dimethoxyphenoxy)aniline 228401-34-1P, 3-(6-Methyl-3-pyridinyloxy)-1-nitrobenzene 228401-35-2P, 3-(6-Methyl-3-pyridinyloxy)aniline 228401-36-3P, 5-Amino-2-(4-methylphenoxy)pyridine Dihydrochloride 228401-37-4P, 4-(3-Thienylthio)-1-nitrobenzene 228401-38-5P, 4-(5-Pyrimidinyloxy)aniline 228401-39-6P, 4-[(2-Methoxy-5-pyridyl)oxy]-1-nitrobenzene 228401-40-9P, 4-(2-Methyl-4-pyridinyloxy)aniline 228401-41-0P, Methyl(4-nitrophenyl)(4-pyridyl)amine 228401-43-2P, 4-(3-Methoxycarbonyl-4-methoxyphenoxy)-1-nitrobenzene 228401-44-3P, 4-(3-Carboxy-4-methoxyphenoxy)-1-nitrobenzene 284462-37-9P, 4-[2-(N-Methylcarbamoyl)-4-pyridyloxy]aniline 284462-38-0P, 5-(4-Nitrophenoxy)isoindoline-1,3-dione 284462-39-1P, 5-(4-Aminophenoxy)isoindoline-1,3-dione 284462-46-0P, 4-[3-(N-Methylcarbamoyl)-4-methoxyphenoxy]-1-nitrobenzene 284462-47-1P, 4-[3-(N-Methylcarbamoyl)-4-methoxyphenoxy]aniline 284462-55-1P, 4-(3-Ethoxycarbonylphenoxy)-1-nitrobenzene 284462-56-2P, 4-(3-N-Methylcarbamoylphenoxy)aniline 284462-78-8P, 3-[2-(N-Methylcarbamoyl)-4-pyridyloxy]aniline 284462-79-9P, 3-(2-Carbamoyl-4-pyridyloxy)aniline 284462-80-2P, 4-(2-Carbamoyl-4-pyridyloxy)aniline 284462-84-6P, 4-(4-Methylsulfonylphenoxy)aniline 432050-13-0P, 4-(3-Thienylthio)aniline 432050-14-1P, 4-[(2-Methoxy-5-pyridyl)oxy]aniline 432050-15-2P, Methyl(4-aminophenyl)(4-pyridyl)amine 432050-16-3P, 4-[Hydroxy(4-pyridyl)methyl]aniline
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(intermediate; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

IT 284461-54-7P, N-[2-Methoxy-5-(trifluoromethyl)phenyl]-N'-[4-(1,3-dioxoisoindolin-5-yloxy)phenyl]urea 284670-98-0P, N,N'-Bis[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea 432050-17-4P 432050-18-5P 432050-19-6P, N,N'-Bis(2-methoxy-3-quinolinyl)urea 432050-20-9P 432050-21-0P, N-[5-Trifluoromethyl-2-pyridyl]-N'-[3-(4-pyridylthio)phenyl]urea 432050-22-1P, N-(2-Methoxy-3-quinolinyl)-N'-[4-(2-(N-Methylcarbamyl)-4-pyridyloxy)phenyl]urea 432050-23-2P, N-(2-Methoxy-3-quinolyl)-N'-[4-[3-(N-methylcarbamoyl)phenoxy]phenyl]urea 432050-24-3P, N-(2-Methoxy-3-quinolyl)-N'-[4-(2-carbamoyl-4-pyridyloxy)phenyl]urea 432050-25-4P, N-(2-Methoxy-3-quinolyl)-N'-[3-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea 432050-26-5P, N-(2-Methoxy-3-quinolyl)-N'-[3-(2-carbamoyl-4-pyridyloxy)phenyl]urea 432050-27-6P, N-(2-Methoxy-3-quinolyl)-N'-[4-[3-(N-isopropylcarbamoyl)phenoxy]phenyl]urea 432050-28-7P, N-(2-Methoxy-3-quinolyl)-N'-[4-[4-methoxy-3-(N-methylcarbamoyl)phenoxy]phenyl]urea 432050-29-8P, N-(3-Isoquinolyl)-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea 432050-30-1P, N-(4-tert-Butyl-2-pyridinyl)-N'-(4-methylphenyl)urea 432050-31-2P, N-(4-tert-Butyl-2-pyridinyl)-N'-(4-fluorophenyl)urea 432050-32-3P, N-(4-tert-Butyl-2-pyridinyl)-N'-(1-naphthyl)urea 432050-33-4P, N-(4-tert-Butyl-2-pyridinyl)-N'-[4-(4-methoxyphenoxy)phenyl]urea 432050-34-5P, N-(5-Trifluoromethyl-2-pyridinyl)-N'-[4-(4-pyridylmethyl)phenyl]urea 432050-35-6P, N-(3-Isoquinolyl)-N'-(4-methylphenyl)urea 432050-36-7P, N-(3-Isoquinolyl)-N'-(4-fluorophenyl)urea 432050-37-8P, N-(3-Isoquinolyl)-N'-(2,3-dichlorophenyl)urea 432050-38-9P, N-(3-Isoquinolyl)-N'-(1-naphthyl)urea 432050-39-0P, N-(3-Isoquinolyl)-N'-[4-(4-pyridinylmethyl)phenyl]urea 432050-40-3P, N-(3-Quinolyl)-N'-[4-(4-pyridinylmethyl)phenyl]urea 432050-41-4P, N-(4-tert-Butyl-2-pyridyl)-N'-(4-(4-methylphenoxy)phenyl)urea 432050-42-5P, N-(4-tert-Butyl-2-pyridyl)-N'-(4-

(4-pyridyloxy)phenyl)urea 432050-43-6P, N-(4-tert-Butyl-2-pyridyl)-N'-(4-(4-pyridinylthio)phenyl)urea 432050-44-7P, N-(4-tert-Butyl-2-pyridyl)-N'-(3-(4-pyridinylthio)phenyl)urea 432050-45-8P 432050-46-9P 432050-47-0P 432050-48-1P 432050-49-2P 432050-50-5P 432050-51-6P, N-(1-(4-Methyl-1-piperazinyl)isoquinol-3-yl)-N'-(4-((4-pyridyl)methyl)phenyl)urea 432050-52-7P, N-(Isoquinol-3-yl)-N'-(4-(3-(methylcarbamoyl)phenoxy)phenyl)urea 432050-53-8P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

IT 75-31-0, Isopropylamine, reactions 86-84-0, 1-Naphthyl isocyanate 98-98-6, Picolinic acid 100-11-8, 4-Nitrobenzyl bromide 100-15-2, N-Methyl-4-nitroaniline 101-77-9, 4,4'-Methylenedianiline 101-79-1, 4-(4-Chlorophenoxy)aniline 106-44-5, 4-Methylphenol, reactions 109-00-2, 3-Hydroxypyridine 123-30-8, 4-Aminophenol 139-59-3, 4-Phenoxyaniline 150-76-5, 4-Methoxyphenol 288-32-4, Imidazole, reactions 350-46-9, 1-Fluoro-4-nitrobenzene 400-74-8, 2-Fluoro-5-nitrobenzotrifluoride 580-17-6, 3-Aminoquinoline 585-79-5, 1-Bromo-3-nitrobenzene 591-27-5, 3-Aminophenol 610-35-5, 4-Hydroxyphthalic acid 620-95-1, 3-Benzylpyridine 622-58-2, 4-Tolyl isocyanate 624-28-2, 2,5-Dibromopyridine 626-61-9, 4-Chloropyridine 626-64-2, 4-Hydroxypyridine 872-31-1, 3-Bromothiophene 1083-48-3, 4-(4-Nitrobenzyl)pyridine 1121-78-4, 5-Hydroxy-2-methylpyridine 1193-02-8, 4-Aminothiophenol 1195-45-5, 4-Fluorophenyl isocyanate 1849-36-1, 4-Nitrothiophenol 2033-89-8, 3,4-Dimethoxyphenol 2103-88-0, 2-Mercapto-4-phenylthiazole 3678-63-5, 4-Chloro-2-methylpyridine 4548-45-2, 2-Chloro-5-nitropyridine 4556-23-4, 4-Mercaptopyridine 4595-59-9, 5-Bromopyrimidine 7379-35-3, 4-Chloropyridine hydrochloride 7781-98-8, Ethyl 3-hydroxybenzoate 16588-75-3, 2-Methoxy-5-(trifluoromethyl)phenyl isocyanate 21101-60-0, 4-(4-Nitrophenylthio)phenol 22948-02-3, 3-Aminothiophenol 24424-99-5, Di-tert-butyl dicarbonate 25267-27-0, Iodobutane 25475-67-6, 3-Aminoisoquinoline 27163-00-4, 4-[(4-Methoxyphenyl)methylamino]aniline 33252-26-5, 2-Amino-4-tert-butylpyridine 36265-31-3, 4-(4-Methylthiophenoxy)-1-nitrobenzene 41195-90-8, 2,3-Dichlorophenyl isocyanate 41295-20-9, 4-(4-Methylphenoxy)aniline 53750-66-6, 4-Chloropyridine-2-carbonyl chloride 73322-01-7, 4-(2-Pyridinylthio)-1-nitrobenzene 74784-70-6, 2-Amino-5-(trifluoromethyl)pyridine 150009-83-9, 3-Amino-2-methoxyquinoline 170893-64-8, 4-(4-Pyridylcarbonyl)aniline 362688-26-4, 1-(4-Methylpiperazinyl)-3-aminoisoquinoline
RL: RCT (Reactant); RACT (Reactant or reagent)
(reactant; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

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AN   2000-499051 [44]    WPIX
CR   2000-499086 [44]; 2002-627516 [67]; 2003-067622 [06]
DNC  C2000-149713
TI   Method of treating a disease mediated by p38 kinase within a host
     comprises administration of urea derivatives e.g. for the treatment of
     cancer or arthritis.
DC   B05
IN   DUMAS, J; KHIRE, U; LOWINGER, T B; MONAHAN, M; NATERO, R; RENICK, J;
     RIEDL, B; SCOTT, W J; SIBLEY, R N; SMITH, R A; WOOD, J E; GUNN, D E;
     HATOUM-MOKDAD, H; NAERO, R; WILLIAM, S J
PA   (FARB) BAYER CORP; (DUMA-I) DUMAS J; (KHIR-I) KHIRE U; (LOWI-I) LOWINGER T
     B; (MONA-I) MONAHAN M; (NAER-I) NAERO R; (RENI-I) RENICK J; (RIED-I) RIEDL
     B; (SIBL-I) SIBLEY R N; (SMIT-I) SMITH R A; (WILL-I) WILLIAM S J; (WOOD-I)
     WOOD J E; (SCOT-I) SCOTT W J
CYC  91
PI   WO 2000041698   A1 20000720 (200044)* EN  148    A61K031-535
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IC ICM A61K031-17; A61K031-506; A61K031-535; A61K031-54; C07D213-63
ICS A61K031-34; A61K031-40; A61K031-4035; A61K031-44; A61K031-445; A61K031-4709; A61K031-4725; A61K031-495; A61K031-497; A61K031-50; A61K031-501; C07D213-70; C07D295-12; C07D295-18

ICA C07C275-36; C07D207-09; C07D209-46; C07D209-48; C07D213-64; C07D213-75; C07D213-81; C07D307-14; C07D401-12

AB WO 200041698 A UPAB: 20040915
NOVELTY - Method of treating a disease mediated by p38 kinase within a host comprises administration of urea derivatives (I) or their salts.

DETAILED DESCRIPTION - Method of treating a disease mediated by p38 kinase within a host comprises administration of urea derivatives of formula (I) or their salts.

D' = NHC(O)NH;

A = L-(M-L1)q which contains up to 40 carbon atoms;

L = 5-6 membered cyclic structure bound to D' containing 0-4 O, N and S, optionally substituted by Y';

L1 = at least 5 membered cyclic group containing 0-4 O, N and S, optionally substituted by L' or Y';

M = bridging group of at least one atom;

q = 1-3;

B' = mono-, di or tricyclic aryl or heteroaryl group which contains up to 30 carbon atoms with at least one 6-membered cyclic group containing 0-4 N, O and S bound directly to D' all optionally substituted by Y';

L' = SO2Rx, C(O)Rx, C(NRy)Rz;

Ry = H or R';

R' = a group containing up to 24 carbon atoms, optionally containing N, S and O optionally substituted by halo;

Rz = H or a group containing up to 30 carbon atoms, optionally containing N, S and O and optionally substituted by halo, OH or R';

Rx = Rz or NRaRb;

Ra, Rb = H, a group containing up to 30 carbon atoms, optionally containing N, S and O and optionally substituted by halo, OH or R', or OSi(Rf)3;

Rf = H or a group containing up to 30 carbon atoms, optionally containing N, S and O and optionally substituted by halo, OH or R'; or

Ra + Rb = 5-7 membered heterocycle containing 1-3 N, S and O, optionally substituted by halo, OH or R'; or

Ra or Rb = C(O) or 1-5C alkylene (optionally substituted with halo, OH or R') bound to L to form an at least 5 membered cyclic structure;

Y' = halo or (W')n;

n = 0-3;

W' = CN, C(O)OR7, C(O)NR7R7, NO2 C(O)R7, OR7, SR7, NR7R7, NR7C(O)OR7, Q-Ar' or a group containing up to 24 carbon atoms, optionally containing N, S and O and optionally substituted by CN, C(O)OR7, C(O)NR7R7, C(O)R7, OR7, SR7, NR7R7, NO2, NR7C(O)R7, NR7C(O)OR7 or halo;

R7 = H or R';

Q = O, S, N(R7), (CH2)m, C(O), CH(OH), (CH2)mO, (CH2)mS, (CH2)mN(R7), O(CH2)m, CHXa, CH(Xa)2, S(CH2)m and NR7(CH2)m;

m = 1-3;

Xa = halo;

Ar' = 5-6 membered aromatic containing 0-2 N, O, S optionally substituted by halo or (Z')n; and

Z' = CN, C(O)OR7, C(O)NR7R7, NO2, OR7, SR7, NR7R7, NR7C(O)OR7 or a group containing up to 24 carbon atoms, optionally containing N, S and O and optionally substituted by CN, C(O)OR7, C(O)NR7R7, C(O)R7, OR7, SR7, NR7R7, NO2, NR7C(O)R7 or NR7C(O)OR7.

ACTIVITY - Cytostatic; osteopathic; antiarthritic; antirheumatic; antibacterial; immunosuppressive; antiarteriosclerotic; neuroprotective; antiinflammatory; gastrointestinal; respiratory; hepatotropic; protozoacide; antidiabetic; cerebroprotective; vulnerary, radioprotective; dermatological.

MECHANISM OF ACTION - P38 kinase inhibitor; tumor necrosis factor alpha (TNF alpha) inhibitor; matrix-destroying metalloprotease (MMP)

inhibitor.

No compound specific biological data given. P38 kinase inhibitory activity was determined using recombinant human p38 (0.5 mu g/ml) mixed with a substrate of myelin basic protein (5 mu g/ml) in kinase buffer and 33P-labeled ATP. The amount of radioactivity incorporated into the substrate was measured. (I) showed p38 IC50 values of 1 nm - 10 mu m.

USE - (I) are used in the treatment of cancerous growth mediated by p38 kinase and in the treatment of rheumatoid arthritis, osteoarthritis, septic arthritis, tumor metastasis, periodontal disease, corneal ulceration, proteinuria, coronary thrombosis from atherosclerotic plaque, aneurysmal aortic birth control, dystrophobic epidermolysis bullosa, degenerative cartilage loss following traumatic joint injury, osteopenias mediated by matrix-destroying metalloprotease (MMP) activity, temperomandibular joint disease, demyelating disease of the nervous system, rheumatic fever, bone reabsorption, postmenopausal osteoporosis, sepsis, gram negative sepsis, septic shock, systemic inflammatory response syndrome, inflammatory bowel syndrome, inflammatory bowel disease, Jarisch-Herxheimer reaction, asthma, adult respiratory distress syndrome, acute pulmonary fibrotic disease, pulmonary sarcoidosis, allergic respiratory disease, silicosis, alveolar injury, hepatic failure, liver disease during acute inflammation, severe alcoholic hepatitis, malaria, non-insulin-dependant diabetes mellitus (NIDDM), congestive heart failure, atherosclerosis, Alzheimer's disease, acute encephalitis, brain injury, multiple sclerosis, lymphoid malignancy, pancreatitis, impaired wound healing in infection, inflammation and cancer, myelodysplastic syndrome, systemic lupus erthematosus, billary cirrhosis, bowel necrosis, psoriasis, radiation injury/toxicity following administration of monoclonal antibodies, host-versus-graft reaction, lung allograft rejection or complications following hip replacement (claimed).

ADVANTAGE - Unlike prior art compositions for treating osteoarthritis, rheumatoid arthritis and septic arthritis p38 kinase inhibitors halt or reverse the progression of cartilage loss and remove or delay the need for surgical intervention. (I) are less toxic than prior art p38 kinase inhibitors.

Dwg.0/0

FS CPI

FA AB; GI; DCN

MC CPI: B06-H; B07-H; B10-A13D; B14-A01; B14-A03B; B14-C03; B14-C09; B14-D07C; B14-E10; B14-F07; B14-F10; B14-G02; B14-H01; B14-J01A; B14-J01B; B14-K01A; B14-K01B; B14-L06; B14-N17A; B14-N17B; B14-N17C; B14-S04

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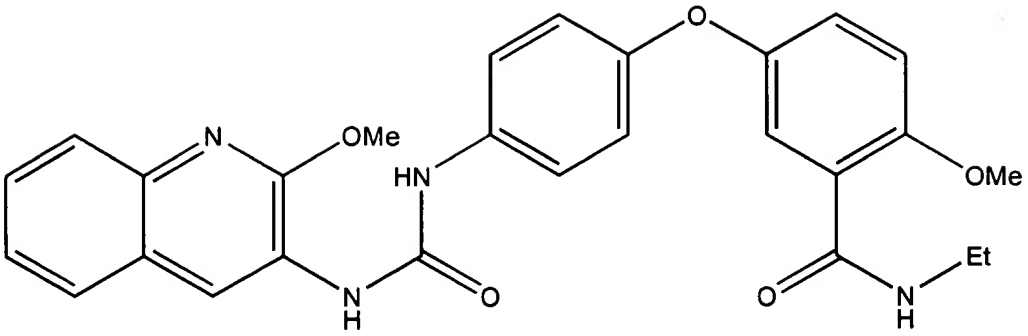

1-(4-(3-(ethylcarbamoyl)-4-methoxyphenoxy)phenyl)-3-(2-methoxyquinolin-3-yl)urea

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* the IDE default display format and the ED field has been added,    *
* effective March 20, 2005.  A new display format, IDERL, is now     *
* available and contains the CA role and document type information.  *
*                                                                    *
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Experimental and calculated property data are now available. For more
information enter HELP PROP at an arrow prompt in the file or refer
to the file summary sheet on the web at:
http://www.cas.org/ONLINE/DBSS/registryss.html

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RN   774552-58-8  REGISTRY
ED   Entered STN:  04 Nov 2004
CN   Urea, N-(4-methylphenyl)-N'-3-quinolinyl- (9CI)  (CA INDEX NAME)
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LC   STN Files:   CHEMCATS
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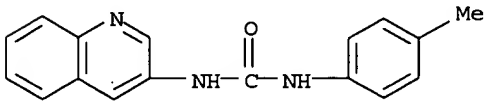


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     (CA INDEX NAME)
OTHER NAMES:
CN   N-(3-Quinolyl)-N'-[4-(4-pyridinylmethyl)phenyl]urea
FS   3D CONCORD
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MF C22 H18 N4 O
SR CA
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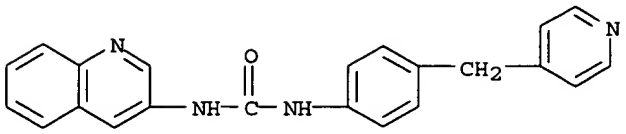

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L55 ANSWER 3 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-39-0 REGISTRY
ED Entered STN: 19 Jun 2002
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OTHER NAMES:
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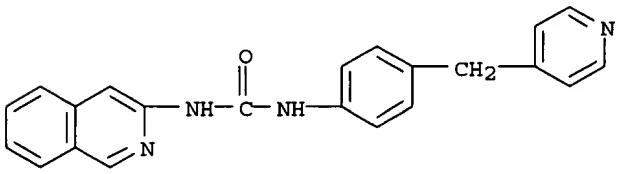

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RN 432050-38-9 REGISTRY
ED Entered STN: 19 Jun 2002
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OTHER NAMES:
CN N-(3-Isoquinolyl)-N'-(1-naphthyl)urea
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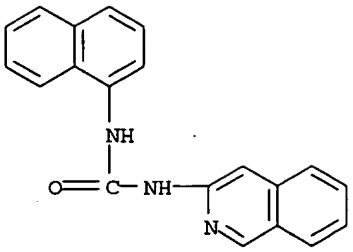

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L55 ANSWER 5 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
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ED Entered STN: 19 Jun 2002
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OTHER NAMES:
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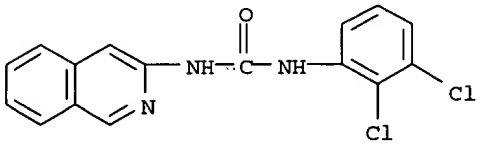

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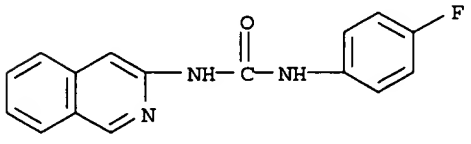
1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

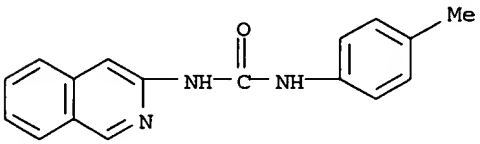
L55 ANSWER 6 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN **432050-36-7** REGISTRY
ED Entered STN: 19 Jun 2002
CN Urea, N-(4-fluorophenyl)-N'-3-isoquinolinyl- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN N-(3-Isoquinolyl)-N'-(4-fluorophenyl)urea
FS 3D CONCORD
MF **C16 H12 F N3 O**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

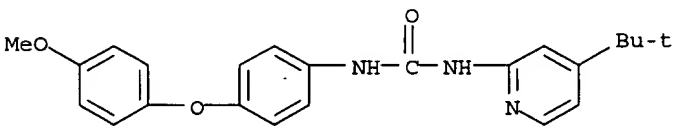
L55 ANSWER 7 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-35-6 REGISTRY
ED Entered STN: 19 Jun 2002
CN Urea, N-3-isoquinolinyl-N'-(4-methylphenyl)- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN N-(3-Isoquinolyl)-N'-(4-methylphenyl)urea
FS 3D CONCORD
MF C17 H15 N3 O
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L55 ANSWER 8 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-33-4 REGISTRY
ED Entered STN: 19 Jun 2002
CN Urea, N-[4-(1,1-dimethylethyl)-2-pyridinyl]-N'-[4-(4-methoxyphenoxy)phenyl]- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN N-(4-tert-Butyl-2-pyridinyl)-N'-[4-(4-methoxyphenoxy)phenyl]urea
FS 3D CONCORD
MF C23 H25 N3 O3
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

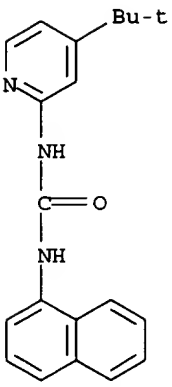


PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L55 ANSWER 9 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-32-3 REGISTRY
ED Entered STN: 19 Jun 2002
CN Urea, N-[4-(1,1-dimethylethyl)-2-pyridinyl]-N'-1-naphthalenyl- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN N-(4-tert-Butyl-2-pyridinyl)-N'-(1-naphthyl)urea
FS 3D CONCORD

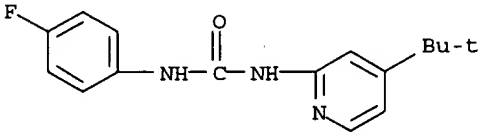
MF **C20 H21 N3 O**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L55 ANSWER 10 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-31-2 REGISTRY
ED Entered STN: 19 Jun 2002
CN **Urea, N-[4-(1,1-dimethylethyl)-2-pyridinyl]-N'-(4-fluorophenyl)-(9CI)** (CA INDEX NAME)
OTHER NAMES:
CN **N-(4-tert-Butyl-2-pyridinyl)-N'-(4-fluorophenyl)urea**
FS 3D CONCORD
MF **C16 H18 F N3 O**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

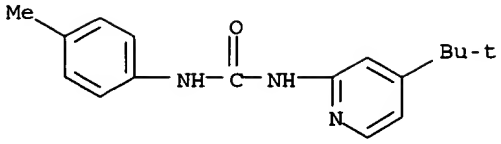


PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L55 ANSWER 11 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN **432050-30-1** REGISTRY
ED Entered STN: 19 Jun 2002
CN **Urea, N-[4-(1,1-dimethylethyl)-2-pyridinyl]-N'-(4-methylphenyl)-(9CI)** (CA INDEX NAME)
OTHER NAMES:
CN N-(4-tert-Butyl-2-pyridinyl)-N'-(4-methylphenyl)urea
FS 3D CONCORD
MF **C17 H21 N3 O**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

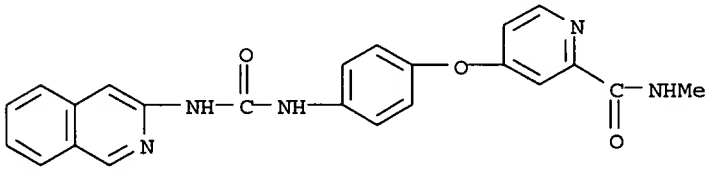
Search done by Noble Jarrell



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

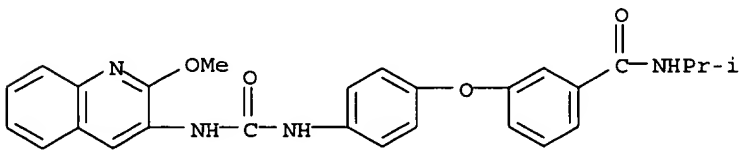
L55 ANSWER 12 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN **432050-29-8** REGISTRY
ED Entered STN: 19 Jun 2002
CN **2-Pyridinecarboxamide, 4-[4-[[(3-isoquinolinylamino)carbonyl]amino]phenoxy]-N-methyl- (9CI)** (CA INDEX NAME)
OTHER NAMES:
CN N-(3-Isoquinolyl)-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea
FS 3D CONCORD
MF **C23 H19 N5 O3**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L55 ANSWER 13 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN **432050-27-6** REGISTRY
ED Entered STN: 19 Jun 2002
CN Benzamide, 3-[4-[[[(2-methoxy-3-quinolinyl)amino]carbonyl]amino]phenoxy]-N-(1-methylethyl)- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN N-(2-Methoxy-3-quinolyl)-N'-[4-[3-(N-isopropylcarbamoyl)phenoxy]phenyl]urea
FS 3D CONCORD
MF **C27 H26 N4 O4**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

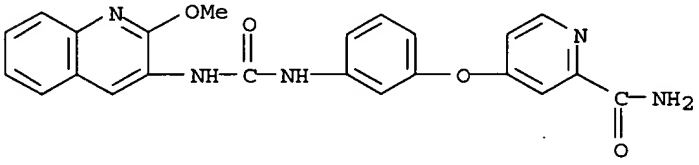


Search done by Noble Jarrell

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

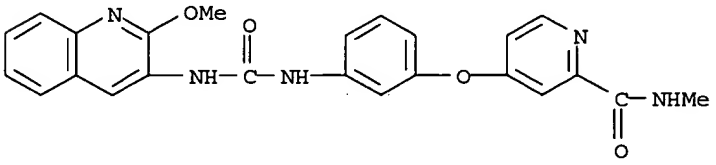
L55 ANSWER 14 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-26-5 REGISTRY
ED Entered STN: 19 Jun 2002
CN **2-Pyridinecarboxamide, 4-[3-[[[(2-methoxy-3-quinolinyl)amino]carbonyl]amino]phenoxy]- (9CI)** (CA INDEX NAME)
OTHER NAMES:
CN N-(2-Methoxy-3-quinolyl)-N'-[3-(2-carbamoyl-4-pyridyloxy)phenyl]urea
FS 3D CONCORD
MF **C23 H19 N5 O4**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L55 ANSWER 15 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-25-4 REGISTRY
ED Entered STN: 19 Jun 2002
CN **2-Pyridinecarboxamide, 4-[3-[[[(2-methoxy-3-quinolinyl)amino]carbonyl]amino]phenoxy]-N-methyl- (9CI)** (CA INDEX NAME)
OTHER NAMES:
CN N-(2-Methoxy-3-quinolyl)-N'-[3-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea
FS 3D CONCORD
MF **C24 H21 N5 O4**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

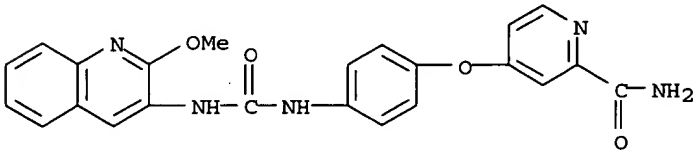


PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L55 ANSWER 16 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN **432050-24-3** REGISTRY
ED Entered STN: 19 Jun 2002
CN **2-Pyridinecarboxamide, 4-[4-[[[(2-methoxy-3-**

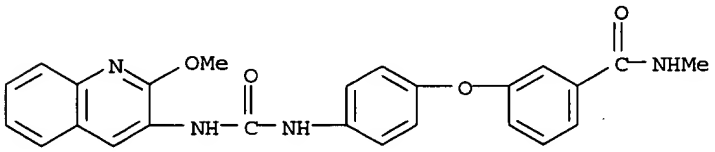
quinolinyl)amino]carbonyl]amino]phenoxy]- (9CI) (CA INDEX NAME)
OTHER NAMES:
CN N-(2-Methoxy-3-quinolyl)-N'-[4-(2-carbamoyl-4-pyridyloxy)phenyl]urea
FS 3D CONCORD
MF **C23 H19 N5 O4**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

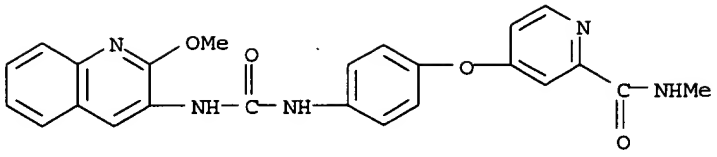
L55 ANSWER 17 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-23-2 REGISTRY
ED Entered STN: 19 Jun 2002
CN **Benzamide, 3-[4-[[[(2-methoxy-3-quinolinyl)amino]carbonyl]amino]phenoxy]-N-methyl- (9CI)** (CA INDEX NAME)
OTHER NAMES:
CN N-(2-Methoxy-3-quinolyl)-N'-[4-[3-(N-methylcarbamoyl)phenoxy]phenyl]urea
FS 3D CONCORD
MF **C25 H22 N4 O4**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

4 REFERENCES IN FILE CA (1907 TO DATE)
4 REFERENCES IN FILE CAPLUS (1907 TO DATE)

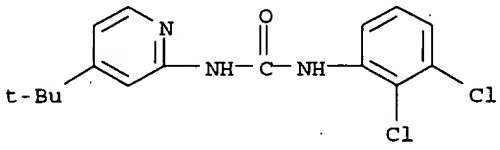
L55 ANSWER 18 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-22-1 REGISTRY
ED Entered STN: 19 Jun 2002
CN **2-Pyridinecarboxamide, 4-[4-[[[(2-methoxy-3-quinolinyl)amino]carbonyl]amino]phenoxy]-N-methyl- (9CI)** (CA INDEX NAME)
OTHER NAMES:
CN N-(2-Methoxy-3-quinolinyl)-N'-[4-(2-(N-Methylcarbamyl)-4-pyridyloxy)phenyl]urea
FS 3D CONCORD
MF **C24 H21 N5 O4**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

5 REFERENCES IN FILE CA (1907 TO DATE)
5 REFERENCES IN FILE CAPLUS (1907 TO DATE)

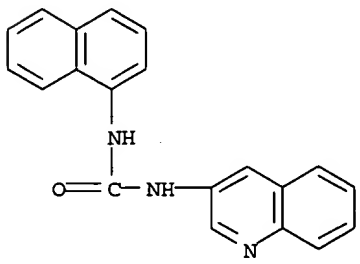
L55 ANSWER 19 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN 432050-17-4 REGISTRY
ED Entered STN: 19 Jun 2002
CN **Urea, N-(2,3-dichlorophenyl)-N'-[4-(1,1-dimethylethyl)-2-pyridinyl]-(9CI)** (CA INDEX NAME)
OTHER NAMES:
CN **N-(4-tert-Butylpyridyl)-N'-(2,3-dichlorophenyl)urea**
FS 3D CONCORD
MF **C16 H17 Cl2 N3 O**
SR CA
LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

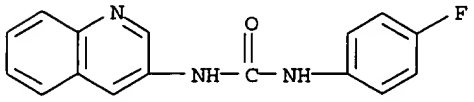
3 REFERENCES IN FILE CA (1907 TO DATE)
3 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L55 ANSWER 20 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN **413613-26-0** REGISTRY
ED Entered STN: 12 May 2002
CN Urea, N-1-naphthalenyl-N'-3-quinolinyl- (9CI) (CA INDEX NAME)
FS 3D CONCORD
MF **C20 H15 N3 O**
SR Chemical Library
Supplier: ChemBridge Corporation
LC STN Files: CHEMCATS



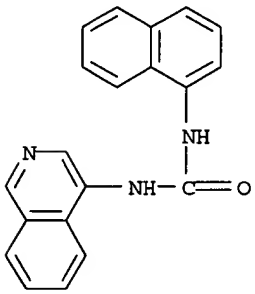
PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L55 ANSWER 21 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN **309284-04-6** REGISTRY
ED Entered STN: 18 Dec 2000
CN Urea, N-(4-fluorophenyl)-N'-3-quinolinyl- (9CI) (CA INDEX NAME)
FS 3D CONCORD
MF **C16 H12 F N3 O**
SR Chemical Library
Supplier: ChemDiv, Inc.
LC STN Files: CHEMCATS



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L55 ANSWER 22 OF 22 REGISTRY COPYRIGHT 2005 ACS on STN
RN **304510-28-9** REGISTRY
ED Entered STN: 27 Nov 2000
CN Urea, N-4-isoquinolinyl-N'-1-naphthalenyl- (9CI) (CA INDEX NAME)
FS 3D CONCORD
MF **C20 H15 N3 O**
SR Chemical Library
Supplier: AsInEx
LC STN Files: CHEMCATS



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

=> d his full

(FILE 'HOME' ENTERED AT 09:19:30 ON 06 JUN 2005)

FILE 'HCAPLUS' ENTERED AT 09:19:43 ON 06 JUN 2005
L1 1 SEA ABB=ON PLU=ON US20020065296/PN OR US99-115878/AP,PRN

FILE 'REGISTRY' ENTERED AT 09:20:28 ON 06 JUN 2005

FILE 'HCAPLUS' ENTERED AT 09:20:29 ON 06 JUN 2005
L2 TRA L1 1- RN : 157 TERMS

FILE 'REGISTRY' ENTERED AT 09:20:30 ON 06 JUN 2005

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L3          157 SEA ABB=ON  PLU=ON  L2

     FILE 'WPIX' ENTERED AT 09:20:35 ON 06 JUN 2005
L4            1 SEA ABB=ON  PLU=ON  US20020065296/PN OR US99-115878/AP,PRN

     FILE 'REGISTRY' ENTERED AT 10:09:52 ON 06 JUN 2005
L5           78 SEA ABB=ON  PLU=ON  L3 AND NC5/ES
L6           13 SEA ABB=ON  PLU=ON  L5 AND NC5-C6/ES
L7            5 SEA ABB=ON  PLU=ON  L3 AND NR=4 AND 2 46.150.18/RID AND
                NC5-C6/ES
L8            0 SEA ABB=ON  PLU=ON  L3 AND N(1A)ETHYL
L9              QUE ABB=ON  PLU=ON  (PMS OR MAN OR IDS)/CI OR COMPD OR
                COMPOUND OR UNSPECIFIED OR (D OR T)/ELS
L10         458 SEA ABB=ON  PLU=ON  C17H21N3O AND NR=2
L11         106 SEA ABB=ON  PLU=ON  L10 AND NC5/ES AND 46.150.18/RID
L12         103 SEA ABB=ON  PLU=ON  L11 NOT L9
L13           7 SEA ABB=ON  PLU=ON  L12 AND DIMETHYLETHYL
                SEL RN 2 L13
L14           1 SEA ABB=ON  PLU=ON  432050-30-1/BI AND L13
L15           9 SEA ABB=ON  PLU=ON  C16H18FN3O AND NR=2 AND NC5/ES AND
                46.150.18/RID
L16           1 SEA ABB=ON  PLU=ON  L15 AND UREA
L17          12 SEA ABB=ON  PLU=ON  C16H17CL2N3O AND NR=2 AND NC5/ES AND
                46.150.18/RID
L18           1 SEA ABB=ON  PLU=ON  L17 AND UREA
L19           3 SEA ABB=ON  PLU=ON  L3 AND C6-C6/ES
L20         111 SEA ABB=ON  PLU=ON  C20H21N3O AND NR=3 AND NC5/ES
L21           3 SEA ABB=ON  PLU=ON  L20 AND C6-C6/ES
L22           1 SEA ABB=ON  PLU=ON  432050-32-3/BI AND L21
L23         488 SEA ABB=ON  PLU=ON  C23H25N3O3 AND NR=3
L24          81 SEA ABB=ON  PLU=ON  L23 AND NC5/ES AND 2 46.150.18/RID
L25          81 SEA ABB=ON  PLU=ON  L24 NOT L9
L26           5 SEA ABB=ON  PLU=ON  L25 AND UREA AND METHOXYPHEN?
                SEL RN 3 L26
L27           1 SEA ABB=ON  PLU=ON  432050-33-4/BI AND L26
L28        1392 SEA ABB=ON  PLU=ON  C17H15N3O NOT L9
L29         124 SEA ABB=ON  PLU=ON  L28 AND NR=3 AND NC5-C6/ES AND 46.150.18/RI
                D
L30          20 SEA ABB=ON  PLU=ON  L29 AND UREA
                D STR TOT
                SEL RN L30 1 10
L31           2 SEA ABB=ON  PLU=ON  (432050-35-6/BI OR 774552-58-8/BI) AND L30
L32          12 SEA ABB=ON  PLU=ON  C16H12FN3O AND NR=3 AND NC5-C6/ES AND
                46.150.18/RID
                SEL RN 4 8 L32
L33           2 SEA ABB=ON  PLU=ON  (309284-04-6/BI OR 432050-36-7/BI) AND L32
L34          24 SEA ABB=ON  PLU=ON  C16H11CL2N3O AND NC5-C6/ES AND 46.150.18/RI
                D
                SEL RN L34 10
L35           1 SEA ABB=ON  PLU=ON  432050-37-8/BI AND L34
L36          17 SEA ABB=ON  PLU=ON  C20H15N3O AND C6-C6/ES AND NC5-C6/ES AND
                NR=4
                D STR TOT
                SEL RN 2 4 5 L36
L37           3 SEA ABB=ON  PLU=ON  (304510-28-9/BI OR 413613-26-0/BI OR
                432050-38-9/BI) AND L36
L38          23 SEA ABB=ON  PLU=ON  C22H18N4O AND NR=4 AND NC5-C6/ES AND
                NC5/ES AND 46.150.18/RID
                D SCA
L39           2 SEA ABB=ON  PLU=ON  L38 AND UREA AND PYRIDINYLMETHYL
                D SCA
                D COS
L40        1130 SEA ABB=ON  PLU=ON  (C25H22N4O4 OR C24H21N5O4 OR C24H21N5O4 OR
                C24H21N5O4 OR C23H19N5O4 OR C27H26N4O4 OR C27H26N4O5 OR
                C23H19N5O3) AND NR=4 NOT L9
L41          60 SEA ABB=ON  PLU=ON  L40 AND NC5-C6/ES AND (2 46.150.18/RID OR
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                NC5/ES AND 46.150.18/RID)
                SEL RN L41 33 29 30
L42           3 SEA ABB=ON  PLU=ON  (432050-24-3/BI OR 432050-27-6/BI OR
                432050-29-8/BI) AND L41
L43           8 SEA ABB=ON  PLU=ON  C25H22N4O4 AND 2 46.150.18/RID AND
                NC5-C6/ES
                D SCA
L44           1 SEA ABB=ON  PLU=ON  L43 AND BENZAMIDE
L45           4 SEA ABB=ON  PLU=ON  C24H21N5O4 AND NR=4 AND NC5-C6/ES AND
                46.150.18/RID AND NC5/ES
L46           2 SEA ABB=ON  PLU=ON  L45 AND PYRIDINECARBOXAMIDE
L47           4 SEA ABB=ON  PLU=ON  C24H21N5O4 AND NC5-C6/ES AND NC5/ES AND
                46.150.18/RID
L48          12 SEA ABB=ON  PLU=ON  L3 AND NC5-C6/ES AND NC5/ES AND 46.150.18/R
                ID AND NR=4
L49           3 SEA ABB=ON  PLU=ON  C23H19N5O4 AND NR=4 AND NC5-C6/ES AND
                NC5/ES AND 46.150.18/RID
L50           2 SEA ABB=ON  PLU=ON  L49 AND PYRIDINECARBOXAMIDE
L51           9 SEA ABB=ON  PLU=ON  C27H26N4O5 AND NR=4 AND 2 46.150.18/RID
                AND NC5-C6/ES
L52         262 SEA ABB=ON  PLU=ON  C23H19N5O3
L53           6 SEA ABB=ON  PLU=ON  L52 AND NR=4 AND NC5/ES AND NC5-C6/ES AND
                46.150.18/RID
L54           1 SEA ABB=ON  PLU=ON  L53 AND PYRIDINECARBOXAMIDE
L55          22 SEA ABB=ON  PLU=ON  L14 OR L16 OR L18 OR L22 OR L27 OR L31 OR
                L33 OR L35 OR L37 OR L39 OR L42 OR L44 OR L46 OR L50 OR L54

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L56           6 SEA ABB=ON  PLU=ON  L55

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L57           0 SEA ABB=ON  PLU=ON  L55

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L58         427 SEA ABB=ON  PLU=ON  ("DUMAS J"/AU OR "DUMAS J B"/AU OR "DUMAS
                J C"/AU OR "DUMAS J F"/AU OR "DUMAS J G"/AU OR "DUMAS J I"/AU
                OR "DUMAS J J"/AU OR "DUMAS J L"/AU OR "DUMAS J M"/AU OR
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                E DUMAS JACQUES/AU
L59         104 SEA ABB=ON  PLU=ON  ("DUMAS JACQUES"/AU OR "DUMAS JACQUES
                P"/AU)
                E RIEDL B/AU
L60         170 SEA ABB=ON  PLU=ON  ("RIEDL B"/AU OR "RIEDL BAUCH VACLAV"/AU
                OR "RIEDL BERNARD"/AU OR "RIEDL BERNARD Y"/AU OR "RIEDL
                BERND"/AU)
                E KHIRE U/AU
L61          43 SEA ABB=ON  PLU=ON  ("KHIRE U R"/AU OR "KHIRE UDAY"/AU OR
                "KHIRE UDAY R"/AU)
                E MOKDAD H/AU
L62           1 SEA ABB=ON  PLU=ON  "MOKDAD H"/AU
                E HATOUM MOKDAD/AU
L63          28 SEA ABB=ON  PLU=ON  ("HATOUM MOKDAD H"/AU OR "HATOUM MOKDAD
                HOLIA"/AU OR "HATOUM MOKDAD HOLIA N"/AU)
                E MONAHAN K/AU
                E MONAHAN M/AU
L64          37 SEA ABB=ON  PLU=ON  ("MONAHAN M"/AU OR "MONAHAN M K"/AU OR
                "MONAHAN MARY K"/AU OR "MONAHAN MARY KATHERINE"/AU OR "MONAHAN
                MARY KATHERINE C"/AU OR "MONAHAN MARY KETHERINE"/AU)
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L65          46 SEA ABB=ON  PLU=ON  ("LOWINGER T B"/AU OR "LOWINGER TIMOTHY"/AU
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                E SCOTT W/AU
L66         158 SEA ABB=ON  PLU=ON  ("SCOTT W"/AU OR "SCOTT W J"/AU OR "SCOTT
                W J JR"/AU OR "SCOTT W J M"/AU OR "SCOTT W J MERLE"/AU OR
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                "SCOTT W JAMES"/AU OR "SCOTT W JOHN H"/AU)
                E SCOTT WILL/AU
L67         225 SEA ABB=ON  PLU=ON  ("SCOTT WILLIAM"/AU OR "SCOTT WILLIAM
                J"/AU OR "SCOTT WILLIAM J JR"/AU OR "SCOTT WILLIAM JAMES"/AU
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                E SMITH R/AU
L68        1107 SEA ABB=ON  PLU=ON  ("SMITH R"/AU OR "SMITH R A"/AU OR "SMITH
                R A A"/AU OR "SMITH R A D"/AU OR "SMITH R A G"/AU OR "SMITH R
                A H"/AU OR "SMITH R A J"/AU OR "SMITH R A K"/AU OR "SMITH R A
                L"/AU OR "SMITH R A W"/AU OR "SMITH R ABBEY"/AU OR "SMITH R
                ANDREW"/AU OR "SMITH R ARNOLD"/AU)
                E SMITH ROGER/AU
L69         293 SEA ABB=ON  PLU=ON  ("SMITH ROGER"/AU OR "SMITH ROGER A"/AU OR
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                ROGER ASTBURY"/AU)
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L70         308 SEA ABB=ON  PLU=ON  ("WOOD J"/AU OR "WOOD J E"/AU OR "WOOD J E
                JR"/AU OR "WOOD J EDWIN"/AU OR "WOOD J EDWIN JR"/AU OR "WOOD J
                ELDRIDGE"/AU)
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                E WOOD JILL/AU
L71          35 SEA ABB=ON  PLU=ON  ("WOOD JILL"/AU OR "WOOD JILL E"/AU OR
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L72       45365 SEA ABB=ON  PLU=ON  BAYER/CS,PA
L73           5 SEA ABB=ON  PLU=ON  L56 AND (L58 OR L59 OR L60 OR L61 OR L62
                OR L63 OR L64 OR L65 OR L66 OR L67 OR L68 OR L69 OR L70 OR
                L71)
L74           5 SEA ABB=ON  PLU=ON  L56 AND L72
L75           5 SEA ABB=ON  PLU=ON  (L73 OR L74)
L76           1 SEA ABB=ON  PLU=ON  L56 NOT L75

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FILE COVERS 1907 - 6 Jun 2005  VOL 142 ISS 24
FILE LAST UPDATED: 5 Jun 2005  (20050605/ED)

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L75  ANSWER 1 OF 5  HCAPLUS  COPYRIGHT 2005 ACS on STN
AN   2003:874973  HCAPLUS
DN   139:364831
ED   Entered STN:  07 Nov 2003
TI   Preparation of quinolyl, isoquinolyl or pyridyl ureas as inhibitors of raf
     kinase using
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IN **Dumas, Jacques; Riedl, Bernd; Khire, Uday; Sibley, Robert N.; Hatoum-Mokdad, Holia; Monahan, Mary-Katherine; Gunn, David E.; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.**
PA **Bayer** Corporation, USA
SO U.S. Pat. Appl. Publ., 26 pp.
CODEN: USXXCO
DT Patent
LA English
IC ICM C07D041-02
ICS A61K031-4709; A61K031-4439
INCL 514307000; 514313000; 546159000; 546143000; 514310000; 514336000; 546268100
CC 27-16 (Heterocyclic Compounds (One Hetero Atom))
Section cross-reference(s): 1, 7
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI US 2003207914	A1	20031106	US 2002-125369	20020419
PRAI US 2001-367376P	P	20010420		

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
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	ICS	A61K031-4709; A61K031-4439
	INCL	514307000; 514313000; 546159000; 546143000; 514310000; 514336000; 546268100
US 2003207914	NCL	514/307.000; 514/313.000; 546/159.000; 546/143.000; 514/310.000; 514/336.000; 546/268.100
	ECLA	C07D213/40B; C07D215/38C; C07D401/12+217+213; C07D401/12+215+213

OS MARPAT 139:364831
AB Urea derivs. of general formula A-NHCONH-B, A'-CONH-B', and A''-NHCONH-B" or pharmaceutically acceptable salts thereof [wherein A = each (un)substituted tert-butylpyridyl, (trifluoromethyl)pyridyl, isopropylpyridyl, 2-methyl-2-butylpyridyl, or 3-methyl-3-pentylpyridyl; A' = each (un)substituted isoquinolinyl or isoquinolinyl; A" = substituted quinolinyl group; B, B' = independently, (un)substituted bridged cyclic structure of up to 30 carbon atoms of the formula -L-(ML1)q (wherein L comprises a cyclic moiety having at least 5 members and is bound directly to D; L1 comprises a cyclic moiety having at least 5 members; M is a bridging group having at least one atom, q is an integer of from 1-3, and each cyclic structure of L and L1 contains 0-4 members of the group consisting of nitrogen, oxygen and sulfur); B" = (un)substituted up to tricyclic aryl or heteroaryl moiety of up to 30 carbon atoms with a cyclic structure bound directly to D containing at least 5 members with 0-4 members of the group consisting of nitrogen, oxygen and sulfur] are prepared These compds. are useful in treating raf-mediated diseases, in particular cancerous cell growth mediated by a raf kinase. All compds. exemplified, e.g. N-(4-tert-Butylpyridyl)-N'-(2,3-dichlorophenyl)urea, displayed IC50 of between 10 nM and 10 μM against ref kinase.
ST quinolylurea prepn raf kinase inhibitor; isoquinolylurea prepn raf kinase inhibitor; pyridylurea prepn raf kinase inhibitor; phenylpyridylurea prepn raf kinase inhibitor; cancer treatment urea prepn
IT Antitumor agents
Neoplasm
(preparation of quinolyl, isoquinolyl or pyridyl ureas as inhibitors of raf kinase)
IT Gene, animal
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(raf, raf-mediated diseases; preparation of quinolyl, isoquinolyl or pyridyl ureas as inhibitors of raf kinase)
IT 6337-24-2P, 1-Methoxy-4-(4-nitrophenoxy)benzene 13472-85-0P, 5-Bromo-2-methoxypyridine 18994-90-6P, 4-(1-Imidazolylmethyl)-1-nitrobenzene 27692-74-6P, 4-(4-Pyridinylmethyl)aniline 28232-34-0P,

5-Nitro-2-(4-methylphenoxy)pyridine 28232-52-2P, 3-(3-Pyridinyloxy)-1-nitrobenzene 31465-36-8P, 4-(4-Methoxyphenoxy)aniline 32361-76-5P, 3-(4-Nitrobenzyl)pyridine 36089-89-1P, 4-(4-Methylsulfonylphenoxy)-1-nitrobenzene 51834-97-0P, 5-Hydroxy-2-methoxypyridine 56643-85-7P, 4-(1-Imidazolylmethyl)aniline 62248-47-9P, 4-(4-Butoxyphenyl)thio-1-nitrobenzene 62248-51-5P, 4-(4-Butoxyphenyl)thioaniline 64064-63-7P, 4-(6-Methyl-3-pyridinyloxy)-1-nitrobenzene 70991-08-1P, 4-(2-Pyridinylthio)aniline 85666-15-5P, 4-(3-Pyridinylmethyl)aniline 92575-23-0P, 3-(4-Pyridinylthio)aniline 116289-71-5P, 3-(3-Pyridinyloxy)aniline 178809-75-1P, 4-[1-Hydroxy-1-(4-pyridyl)methyl]-1-nitrobenzene 220000-87-3P, 2-(N-Methylcarbamoyl)-4-chloropyridine 228401-26-1P, 3-(Trifluoromethyl)-4-(4-pyridinylthio)nitrobenzene 228401-27-2P, 3-(Trifluoromethyl)-4-(4-pyridinylthio)aniline 228401-28-3P, 4-(4-Phenyl-2-thiazolylthio)-1-nitrobenzene 228401-29-4P, 4-(4-Phenyl-2-thiazolyl)thioaniline 228401-31-8P, 4-(6-Methyl-3-pyridinyloxy)aniline 228401-32-9P, 4-(3,4-Dimethoxyphenoxy)-1-nitrobenzene 228401-33-0P, 4-(3,4-Dimethoxyphenoxy)aniline 228401-36-3P, 5-Amino-2-(4-methylphenoxy)pyridine dihydrochloride 228401-37-4P, 4-(3-Thienylthio)-1-nitrobenzene 228401-38-5P, 4-(5-Pyrimidinyloxy)aniline 228401-39-6P, 4-(2-Methoxy-5-pyridyloxy)-1-nitrobenzene 228401-40-9P, 4-(2-Methyl-4-pyridinyloxy)aniline 228401-41-0P, Methyl(4-nitrophenyl)-4-pyridylamine 228401-43-2P, 4-(3-Methoxycarbonyl-4-methoxyphenoxy)-1-nitrobenzene 228401-44-3P, 4-(3-Carboxy-4-methoxyphenoxy)-1-nitrobenzene 229003-17-2P, 3-(5-Methyl-3-pyridinyloxy)-1-nitrobenzene 284462-84-6P, 4-(4-Methylsulfonylphenoxy)aniline 432050-13-0P, 4-(3-Thienylthio)aniline 432050-14-1P, 4-(2-Methoxy-5-pyridyloxy)aniline 432050-15-2P, Methyl(4-aminophenyl)-4-pyridylamine 432050-16-3P, 4-[1-Hydroxy-1-(4-pyridyl)methyl]aniline 473915-53-6P, 4-(4-tert-Butoxycarbamoylbenzyl)aniline 620625-96-9P, 4-[[2-(Methylcarbamoyl)-3-pyridyl]oxy]aniline
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(intermediate; preparation of quinolyl, isoquinolyl or pyridyl ureas as inhibitors of raf kinase)

IT 139691-76-2, Raf Kinase
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(preparation of quinolyl, isoquinolyl or pyridyl ureas as inhibitors of raf kinase)

IT **432050-17-4P**, N-(4-tert-Butylpyridyl)-N'-(2,3-dichlorophenyl)urea 432050-18-5P, N-(4-tert-Butylpyridyl)-N'-[4-(4-pyridinylmethyl)phenyl]urea 432050-19-6P, N,N'-Bis(2-methoxy-3-quinolinyl)urea] 432050-20-9P, N-(4-tert-Butylpyridyl)-N'-[4-(4-chlorophenoxy)phenyl]urea 432050-21-0P, N-(5-Trifluoromethyl-2-pyridyl)-N'-[3-(4-pyridylthio)phenyl]urea **432050-22-1P**, N-(2-Methoxy-3-quinolinyl)-N'-[4-[2-(N-Methylcarbamyl)-4-pyridyloxy]phenyl]urea 432050-41-4P 432050-42-5P 432050-43-6P 432050-44-7P 432050-45-8P 432050-46-9P 432050-47-0P 432050-48-1P 432050-49-2P 473915-54-7P 473915-55-8P 473915-56-9P, N-[1-(4-Methylpiperazinyl)-3-isoquinolinyl]-N'-[4-(4-pyridinyloxy)phenyl]urea 620625-86-7P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of quinolyl, isoquinolyl or pyridyl ureas as inhibitors of raf kinase)

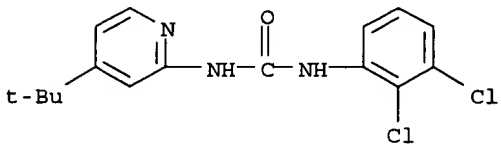
IT 75-44-5, Phosgene 100-11-8, 4-Nitrobenzyl bromide 100-15-2, N-Methyl-4-nitroaniline 101-77-9, 4,4'-Methylenedianiline 101-79-1, 4-(4-Chlorophenoxy)aniline 106-44-5, 4-Methylphenol, reactions 109-00-2, 3-Hydroxypyridine 123-30-8, 4-Aminophenol 139-59-3, 4-Phenoxyaniline 150-76-5, 4-Methoxyphenol 288-32-4, Imidazole, reactions 350-46-9, 1-Fluoro-4-nitrobenzene 400-74-8, 2-Fluoro-5-nitrobenzotrifluoride 530-62-1, N,N'-Carbonyldiimidazole 585-79-5, 1-Bromo-3-nitrobenzene 620-95-1, 3-Benzylpyridine 624-28-2, 2,5-Dibromopyridine 626-61-9, 4-Chloropyridine 626-64-2, 4-Hydroxypyridine 673-09-6, 4-(4-Pyridylthio)aniline 872-31-1,

3-Bromothiophene 1083-48-3, 4-(4-Nitrobenzyl)pyridine 1121-78-4, 5-Hydroxy-2-methylpyridine 1193-02-8, 4-Aminothiophenol 1849-36-1, 4-Nitrothiophenol 2033-89-8, 3,4-Dimethoxyphenol 2103-88-0, 2-Mercapto-4-phenylthiazole 3678-63-5, 4-Chloro-2-picoline 4548-45-2, 2-Chloro-5-nitropyridine 4556-23-4, 4-Mercaptopyridine 4595-59-9, 5-Bromopyrimidine 7379-35-3, 4-Chloropyridine hydrochloride 21101-60-0, 4-(4-Nitrophenylthio)phenol 22948-02-3, 3-Aminothiophenol 24424-99-5, Di-tert-butyl dicarbonate 25267-27-0, Iodobutane 25475-67-6, 3-Aminoisoquinoline 27163-00-4, 4-[(4-Methoxyphenyl)methylamino]aniline 29264-35-5, 4-(3-Carboxy-4-hydroxyphenoxy)-1-nitrobenzene 33252-26-5, 2-Amino-4-tert-butylpyridine 36265-31-3, 4-(4-Methylthiophenoxy)-1-nitrobenzene 41195-90-8, 2,3-Dichlorophenyl isocyanate 41295-20-9, 4-(4-Methylphenoxy)aniline 42732-49-0, 3-Hydroxy-5-methylpyridine 73322-01-7, 4-(2-Pyridinylthio)-1-nitrobenzene 74784-70-6, 2-Amino-5-(trifluoromethyl)pyridine 102877-78-1, 4-(4-Aminophenoxy)pyridine 150009-83-9, 3-Amino-2-methoxyquinoline 170893-64-8, 4-(4-Pyridylcarbonyl)aniline 284462-37-9 362688-26-4, 1-(4-Methylpiperazinyl)-3-aminoisoquinoline
RL: RCT (Reactant); RACT (Reactant or reagent)
(reactant; preparation of quinolyl, isoquinolyl or pyridyl ureas as inhibitors of raf kinase)

IT **432050-17-4P**, N-(4-tert-Butylpyridyl)-N'-(2,3-dichlorophenyl)urea
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of quinolyl, isoquinolyl or pyridyl ureas as inhibitors of raf kinase)

RN 432050-17-4 HCAPLUS

CN Urea, N-(2,3-dichlorophenyl)-N'-[4-(1,1-dimethylethyl)-2-pyridinyl]- (9CI) (CA INDEX NAME)



L75 ANSWER 2 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2002:850357 HCAPLUS

DN 137:352907

ED Entered STN: 08 Nov 2002

TI Preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase for the treatment of tumors and/or cancerous cell growth

IN **Dumas, Jacques**; **Riedl, Bernd**; **Khire, Uday**; **Wood, Jill E.**; Robert, Sibley N.; **Monahan, Mary-Katherine**; Renick, Joel; Gunn, David E.; **Lowinger, Timothy B.**; **Scott, William J.**; **Smith, Roger A.**

PA **Bayer** Corporation, USA

SO U.S. Pat. Appl. Publ., 63 pp., Cont.-in-part of U.S. Ser. No. 758,548.
CODEN: USXXCO

DT Patent

LA English

IC ICM C07D217-22
ICS C07D215-38

INCL 546143000

CC 27-17 (Heterocyclic Compounds (One Hetero Atom))
Section cross-reference(s): 1

FAN.CNT 5

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PI	US 2002165394	A1	20021107	US 2001-777920	20010207
	ZA 2001005751	A	20030714	ZA 2001-5751	20010712

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     US 2002137774         A1     20020926    US 2001-907970          20010719
     WO 2002062763         A2     20020815    WO 2002-US3361          20020207
     WO 2002062763         A3     20021010
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             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL,
             PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG,
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         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
             CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
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PRAI US 1999-115877P       P      19990113
     US 1999-257266        B2     19990225
     US 1999-425228        B2     19991022
     US 2001-758548        A2     20010112
     US 1999-115878P       P      19990113
     US 2001-777920        A      20010207
     US 2001-948915        A1     20010910
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                  INCL    546143000
 US 2002165394    NCL     546/143.000; 546/159.000; 546/306.000; 546/014.000
                  ECLA    C07C275/28; C07C275/30; C07C275/32; C07C275/36;
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                          C07D213/75B2; C07D213/75D3; C07D213/81E; C07D213/82H;
                          C07D215/38C; C07D295/12A1; C07D295/12B1D4;
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                          C07D295/12B1D4; C07D295/18B2D
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                          C07D401/12+215+213; C07D401/12+215+209;
                          C07D401/12+217+213; C07D401/12+213+207; C07C275/40;
                          C07C311/29; C07C317/22; C07D213/75B2; C07D213/82H;
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OS   MARPAT 137:352907
GI
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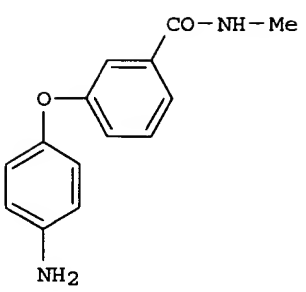

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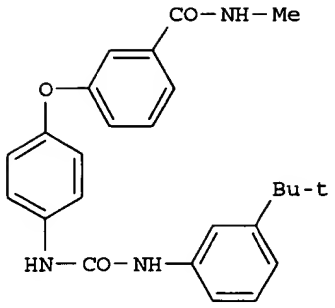


III

AB Title compds. B-NHCONH-L-(M-L1)q (I) [B = (un)substituted pyridyl, quinolinyl, isoquinolinyl; L = 5 or 6 membered cyclic structure; L1 = substituted cyclic moiety having at least 5 members; M = bridging group having at least one atom; q = 1-3; with proviso that L and L1 contain 0-4 hetero atoms, e.g., N, O and S] and their pharmaceutically acceptable salts were prepared For example, coupling of aniline II, e.g., prepared from Et 3-hydroxybenzoate in 4-steps, with bis(trichloromethyl)carbonate followed by 3-tert-butylaniline afforded urea III. In in vitro raf kinase assays, 112-specific examples of compds. I inhibited kinase activity with IC50 values ranging from 10 nM-10 μM. Compds. I are useful for the treatment of cancerous cell growth mediated by raf kinase.

ST quinoline urea prepn inhibition raf kinase antitumor; isoquinoline urea prepn inhibition raf kinase antitumor; pyridine urea prepn inhibition raf kinase antitumor

IT Antitumor agents
Combinatorial chemistry
Human
Neoplasm
Solid phase synthesis
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT 228418-48-2P 284461-33-2P 284461-34-3P 284461-35-4P 284461-36-5P
284461-37-6P 284461-38-7P 284461-39-8P 284461-40-1P 284461-41-2P
284461-42-3P 284461-43-4P 284461-44-5P 284461-45-6P 284461-46-7P
284461-47-8P 284461-48-9P 284461-49-0P 284461-50-3P 284461-51-4P
284461-52-5P 284461-53-6P 284461-54-7P 284461-55-8P 284461-56-9P
284461-57-0P 284461-58-1P 284461-59-2P 284461-60-5P 284461-61-6P
284461-62-7P 284461-63-8P 284461-64-9P 284461-65-0P 284461-66-1P
284461-67-2P 284461-68-3P 284461-69-4P 284461-70-7P 284461-71-8P
284461-72-9P 284461-73-0P 284461-74-1P 284461-75-2P 284461-76-3P
284461-77-4P 284461-78-5P 284461-79-6P 284461-80-9P 284461-81-0P
284461-82-1P 284461-84-3P 284461-85-4P 284461-86-5P 284461-88-7P
284461-89-8P 284461-90-1P 284461-91-2P 284461-92-3P 284461-93-4P
284461-94-5P 284461-95-6P 284461-96-7P 284461-97-8P 284461-98-9P
284462-00-6P 284462-01-7P 284462-02-8P 284462-03-9P 284462-04-0P
284462-05-1P 284462-07-3P 284462-08-4P 284462-09-5P 284462-10-8P
284462-11-9P 284462-12-0P 284462-13-1P 284462-15-3P 284462-16-4P
284462-17-5P 284462-18-6P 284462-19-7P 284462-20-0P 284462-21-1P
284462-22-2P 284462-23-3P 284462-24-4P 284462-25-5P 284462-26-6P
284462-27-7P 284462-28-8P 284462-29-9P 284462-30-2P 284462-31-3P
284462-32-4P 284462-34-6P 284462-35-7P 284462-70-0P 284670-98-0P
432050-20-9P, N-(4-tert-Butylpyridyl)-N'-(4-(4-chlorophenoxy)phenyl) Urea
432050-22-1P 432050-23-2P 432050-24-3P
432050-25-4P 432050-26-5P 432050-27-6P
432050-28-7P 432050-52-7P 447457-08-1P 447457-09-2P 474642-44-9P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(drug candidate; preparation of quinolyl, isoquinolyl or pyridyl-ureas as

inhibitors of raf kinase)

IT 883-62-5P, 3-Methoxy-2-naphthoic acid 13041-60-6P, Methyl 3-methoxy-2-naphthoate 27237-21-4P, 4-(3-Carboxyphenoxy)-1-nitrobenzene 36089-89-1P, 4-(4-Methylsulfonylphenoxy)-1-nitrobenzene 41513-02-4P, 4-Bromo-3-(trifluoromethyl)phenyl isocyanate 50727-06-5P, 5-Hydroxyisoindoline-1,3-dione 51727-15-2P, 4-Chloropyridine-2-carbonyl chloride hydrochloride 54579-63-4P, 4-(3-Carboxyphenoxy)aniline 64064-63-7P 67291-63-8P, 2-Amino-3-methoxynaphthalene 71708-64-0P, 4-(3-(N-Methylcarbamoyl)phenoxy)-1-nitrobenzene 77992-50-8P 119431-22-0P, 3-Chloro-4-(2,2,2-trifluoroacetylamino)phenol 153435-79-1P, N-Methyl-3-bromobenzenesulfonamide 176977-85-8P, Methyl 4-chloropyridine-2-carboxylate hydrochloride 220000-87-3P, 4-Chloro-N-methyl-2-pyridinecarboxamide 228401-15-8P 228401-43-2P, 4-(3-Methoxycarbonyl-4-methoxyphenoxy)-1-nitrobenzene 228401-44-3P, 4-(3-Carboxy-4-methoxyphenoxy)-1-nitrobenzene 252061-66-8P, 5-Hydroxyisoindolin-1-one 284461-99-0P 284462-37-9P, 4-(2-(N-Methylcarbamoyl)-4-pyridyloxy)aniline 284462-38-0P, 5-(4-Nitrophenoxy)isoindoline-1,3-dione 284462-39-1P, 5-(4-Aminophenoxy)isoindoline-1,3-dione 284462-40-4P, 1-(4-tert-Butyl-2-nitrophenyl)-2,5-dimethylpyrrole 284462-41-5P, 5-tert-Butyl-2-(2,5-dimethylpyrrolyl)aniline 284462-42-6P, 4-(2-(N-Methylcarbamoyl)-4-pyridyloxy)-2-methylaniline hydrochloride 284462-43-7P 284462-44-8P, 4-(2-(N-Methylcarbamoyl)-4-pyridyloxy)-2-chloroaniline 284462-45-9P, 4-Chloro-2-methoxy-5-(trifluoromethyl)aniline 284462-46-0P 284462-47-1P 284462-48-2P, 5-(4-Nitrophenoxy)-2-methylisoindoline-1,3-dione 284462-49-3P, 5-(4-Aminophenoxy)-2-methylisoindoline-1,3-dione 284462-51-7P, 4-Chloro-2-(N-(2-morpholin-4-ylethyl)carbamoyl)pyridine 284462-52-8P 284462-53-9P, 4-(1-Oxoisoindolin-5-yloxy)-1-nitrobenzene 284462-54-0P, 4-(1-Oxoisoindolin-5-yloxy)aniline 284462-55-1P, 4-(3-Ethoxycarbonylphenoxy)-1-nitrobenzene 284462-56-2P 284462-57-3P 284462-58-4P 284462-59-5P, 4-(3-(N-Methylsulfamoyl)phenyloxy)benzene 284462-60-8P, 4-(3-(N-Methylsulfamoyl)phenyloxy)-1-nitrobenzene 284462-61-9P, 4-(3-(N-Methylsulfamoyl)phenyloxy)aniline 284462-62-0P 284462-63-1P, 4-Chloro-N-(2-triisopropylsilyloxy)ethylpyridine-2-carboxamide 284462-64-2P 284462-65-3P 284462-66-4P 284462-67-5P, N-(4-Chloro-3-(trifluoromethyl)phenyl)-N'-(4-aminophenyl) Urea 284462-68-6P, N-(4-Chloro-3-(trifluoromethyl)phenyl)-N'-(4-ethoxycarbonylphenyl) Urea 284462-71-1P 284462-97-1P 474642-51-8P, N-(4-Chloro-3-(trifluoromethyl)phenyl)-N'-(3-carboxyphenyl) Urea
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(intermediate; preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT 50-21-5, Lactic acid, reactions 64-19-7, Acetic acid, reactions 65-85-0, Benzoic acid, reactions 69-72-7, Salicylic acid, reactions 75-75-2, Methanesulfonic acid 76-05-1, Trifluoroacetic acid, reactions 77-92-9, Citric acid, reactions 85-47-2, 1-Naphthalenesulfonic acid 87-69-4, Tartaric acid, reactions 90-64-2, Mandelic acid 98-11-3, Benzenesulfonic acid, reactions 103-82-2, Phenylacetic acid, reactions 104-15-4, reactions 110-15-6, Succinic acid, reactions 110-16-7, Maleic acid, reactions 110-17-8, Fumaric acid, reactions 120-18-3, 2-Naphthalenesulfonic acid 144-62-7, Oxalic acid, reactions 1493-13-6, Trifluoromethylsulfonic acid 6915-15-7, Malic acid 7647-01-0, Hydrochloric acid, reactions 7664-38-2, Phosphoric acid, reactions 7664-93-9, Sulphuric acid, reactions 10035-10-6, Hydrobromic acid, reactions
RL: RGT (Reagent); RACT (Reactant or reagent)
(pharmaceutical salt; preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT 139691-76-2, Raf kinase
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

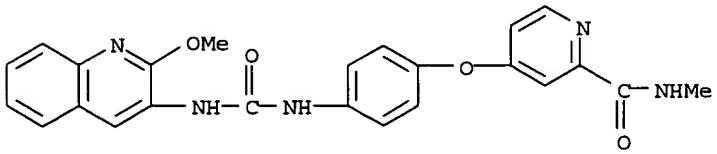
IT 75-31-0, Isopropylamine, reactions 98-58-8, 4-Bromobenzenesulfonyl chloride 98-98-6, Picolinic acid 99-93-4, p-Hydroxyacetophenone

99-98-9, 4-(Dimethylamino)aniline 100-51-6, Benzyl alcohol, reactions
101-79-1, 4-(4-Chlorophenoxy)aniline 106-50-3, p-Phenylenediamine,
reactions 108-00-9, N,N-Dimethylethylenediamine 108-95-2, Phenol,
reactions 109-85-3, 2-Methoxyethylamine 110-13-4, Acetonylacetone
123-30-8, 4-Aminophenol 123-39-7, N-Methylformamide 320-51-4,
4-Chloro-3-(trifluoromethyl)aniline 327-78-6, 4-Chloro-3-
(trifluoromethyl)phenyl isocyanate 349-65-5, 2-Methoxy-5-
(trifluoromethyl)aniline 350-46-9, 1-Fluoro-4-nitrobenzene 371-40-4,
4-Fluoroaniline 393-36-2, 4-Bromo-3-(trifluoromethyl)aniline 407-25-0,
Trifluoroacetic anhydride 462-08-8, 3-Aminopyridine 503-38-8,
Trichloromethyl chloroformate 610-35-5, 4-Hydroxyphthalic acid
619-08-9, 2-Chloro-4-nitrophenol 626-61-9, 4-Chloropyridine 883-99-8,
Methyl 3-hydroxy-2-naphthoate 1121-78-4, 5-Hydroxy-2-methylpyridine
1193-02-8, 4-Aminothiophenol 1664-40-0, N-Phenylethylenediamine
1877-71-0, Mono-methyl isophthalate 2038-03-1, 4-(2-
Aminoethyl)morpholine 2252-63-3, N-(4-Fluorophenyl)piperazine
2524-67-6, 4-Morpholinoaniline 2835-95-2, 5-Amino-2-methylphenol
2835-99-6, 4-Amino-3-methylphenol 2905-24-0, 3-Bromobenzenesulfonyl
chloride 3535-88-4, 5-Tert-Butyl-2-methoxyaniline 3964-52-1,
4-Amino-2-chlorophenol 4548-45-2, 2-Chloro-5-nitropyridine 4795-29-3,
Tetrahydrofurfurylamine 5369-19-7, 3-tert-Butylaniline 6310-19-6,
2-Nitro-4-tert-butylaniline 6628-77-9, 5-Amino-2-methoxypyridine
6927-86-2 7781-98-8, Ethyl 3-hydroxybenzoate 13154-24-0,
Triisopropylsilyl chloride 16588-75-3, 2-Methoxy-5-
(trifluoromethyl)phenyl isocyanate 22948-02-3, 3-Aminothiophenol
25900-61-2, 3-Methylcarbamoylaniline 26116-12-1, 2-Aminomethyl-1-
ethylpyrrolidine 27578-60-5, 1-(2-Aminoethyl)piperidine 30766-22-4,
Methyl 5-hydroxynicotinate 30806-83-8, Ethyl 4-isocyanatobenzoate
33252-26-5, 4-tert-Butyl-2-aminopyridine 34803-66-2,
N-(2-Pyridyl)piperazine 36265-31-3, 4-(4-Methylthiophenoxy)-1-
nitrobenzene 51639-48-6, N-(4-Acetylphenyl)piperazine 106164-64-1
150009-83-9, 3-Amino-2-methoxyquinoline 284462-72-2 284462-73-3,
4-Chloro-N-(2-hydroxyethyl)pyridine-2-carboxamide 284462-74-4,
4-(2-(N-Methylcarbamoyl)-4-pyridyloxy)-2-methylaniline 474642-55-2
RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT 29264-35-5P, 4-(3-Carboxy-4-hydroxyphenoxy)-1-nitrobenzene
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT **432050-22-1P**
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(drug candidate; preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

RN 432050-22-1 HCAPLUS
CN 2-Pyridinecarboxamide, 4-[4-[[[(2-methoxy-3-quinolinyl)amino]carbonyl]amino]phenoxy]-N-methyl- (9CI) (CA INDEX NAME)



L75 ANSWER 3 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN
AN 2002:832761 HCAPLUS
DN 137:337791
ED Entered STN: 01 Nov 2002

TI Preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase

IN **Dumas, Jacques; Riedl, Bernd; Khire, Uday; Sibley, Robert N.; Hatoum-Mokdad, Holia; Monahan, Mary-Katherine; Gunn, David E.; Lowinger, Timothy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.**

PA **Bayer Corporation, USA**

SO PCT Int. Appl., 65 pp.
CODEN: PIXXD2

DT Patent

LA English

IC ICM C07D213-40
ICS C07D401-12; C07D215-38; A61K031-44; A61K031-4709; A61K031-4725; A61K031-47; A61P043-00

CC 27-17 (Heterocyclic Compounds (One Hetero Atom))
Section cross-reference(s): 1, 63

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI WO 2002085857	A2	20021031	WO 2002-US12066	20020418
WO 2002085857	A3	20030116		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CA 2443950	AA	20021031	CA 2002-2443950	20020418
EP 1379505	A2	20040114	EP 2002-725710	20020418
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
JP 2005501813	T2	20050120	JP 2002-583384	20020418
PRAI US 2001-838285	A	20010420		
WO 2002-US12066	W	20020418		

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 2002085857	ICM	C07D213-40
	ICS	C07D401-12; C07D215-38; A61K031-44; A61K031-4709; A61K031-4725; A61K031-47; A61P043-00
WO 2002085857	ECLA	C07D213/40B; C07D215/38C; C07D401/12+215+213; C07D401/12+217+213
JP 2005501813	FTERM	4C031/AA01; 4C031/DA02; 4C031/EA02; 4C031/JA02; 4C055/AA01; 4C055/BA02; 4C055/BA28; 4C055/BB17; 4C055/CA01; 4C055/DA06; 4C055/EA01; 4C063/AA01; 4C063/BB08; 4C063/BB09; 4C063/CC14; 4C063/CC15; 4C063/DD12; 4C063/EE01; 4C086/AA01; 4C086/AA02; 4C086/AA03; 4C086/BC17; 4C086/BC28; 4C086/BC30; 4C086/GA08; 4C086/MA04; 4C086/MA22; 4C086/MA23; 4C086/MA35; 4C086/MA52; 4C086/MA55; 4C086/MA63; 4C086/NA14; 4C086/ZB26; 4C086/ZC20

OS MARPAT 137:337791

AB Title compds. A-D-B (I) [D = NHCONH; A = (un)substituted t-butylpyridyl, etc.; B = (un)substituted bridged cyclic structure, etc.] and analogs were prepared For instance, 4-tert-butyl-2-aminopyridine was coupled to 4-(4-pyridylmethyl)aniline (CH_2Cl_2, CDI, 0°) to give N-(4-tert-butylpyridyl)-N'-[4-(4-pyridinylmethyl)phenyl]urea as a white solid. Example compds. had IC50 between 10nM and 10μM for raf kinase. I are useful for the treatment of cancerous cell growth mediated by raf kinase.

ST inhibition raf kinase quinoline isoquinoline pyridine ureas prepn

IT Antitumor agents

Human
Neoplasm
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT 139691-76-2, Raf kinase
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT **432050-17-4P**, N-(4-tert-Butylpyridyl)-N'-(2,3-dichlorophenyl)urea
432050-18-5P, N-(4-tert-Butylpyridyl)-N'-[4-(4-pyridinylmethyl)phenyl]urea
432050-19-6P, N,N'-[2-Methoxyquinolin-3-yl]urea 432050-20-9P,
N-(4-tert-Butylpyridyl)-N'-[4-(4-chlorophenoxy)phenyl]urea 432050-21-0P,
N-[5-(Trifluoromethyl)pyridin-2-yl]-N'-[3-(4-pyridylthio)phenyl]urea
432050-22-1P 432050-41-4P 432050-42-5P 432050-43-6P
432050-44-7P 432050-45-8P 432050-46-9P 432050-47-0P 432050-48-1P
432050-49-2P 473915-54-7P 473915-55-8P 473915-56-9P 473915-57-0P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT 100-11-8, 4-Nitrobenzyl bromide 100-15-2, N-Methyl-4-nitroaniline
101-77-9, 4,4'-Methylenedianiline 101-79-1, 4-(4-Chlorophenoxy)aniline
106-44-5, 4-Methylphenol, reactions 109-00-2, 3-Hydroxypyridine
123-30-8, 4-Aminophenol 123-39-7, N-Methylformamide 139-59-3,
4-Phenoxyaniline 150-76-5, 4-Methoxyphenol 288-32-4, Imidazole,
reactions 350-46-9, 1-Fluoro-4-nitrobenzene 400-74-8,
2-Fluoro-5-nitrobenzotrifluoride 585-79-5, 1-Bromo-3-nitrobenzene
620-95-1, 3-Benzylpyridine 624-28-2, 2,5-Dibromopyridine 626-61-9,
4-Chloropyridine 673-09-6, 4-(4-Pyridylthio)aniline 872-31-1,
3-Bromothiophene 1083-48-3, 4-(4-Nitrobenzyl)pyridine 1121-78-4,
5-Hydroxy-2-methylpyridine 1849-36-1, 4-Nitrothiophenol 2033-89-8,
3,4-Dimethoxyphenol 2103-88-0, 2-Mercapto-4-phenylthiazole 3678-63-5
4548-45-2, 2-Chloro-5-nitropyridine 4556-23-4, 4-Mercaptopyridine
4595-59-9, 5-Bromopyrimidine 7379-35-3, 4-Chloropyridine hydrochloride
21101-60-0, 4-(4-Nitrophenylthio)phenol 22948-02-3, 3-Aminothiophenol
25267-27-0, Iodobutane 25475-67-6, 3-Aminoisoquinoline 27163-00-4,
4-[(4-Methoxyphenyl)methylamino]aniline 29264-35-5, 4-(3-Carboxy-4-hydroxyphenoxy)-1-nitrobenzene 33252-26-5, 2-Amino-4-tert-butylpyridine
36265-31-3, 4-(4-Methylthiophenoxy)-1-nitrobenzene 41195-90-8,
2,3-Dichlorophenyl isocyanate 41295-20-9, 4-(4-Methylphenoxy)aniline
42732-49-0, 3-Hydroxy-5-methylpyridine 73322-01-7, 4-(2-Pyridinylthio)-1-nitrobenzene 74784-70-6, 2-Amino-5-(trifluoromethyl)pyridine
102877-78-1 150009-83-9, 3-Amino-2-methoxyquinoline 170893-64-8,
4-(4-Pyridylcarbonyl)aniline 284462-37-9, 4-((2-(Methylcarbamoyl)pyridin-4-yl)oxy)aniline 361551-95-3, 3-(4-Pyridylmethyl)aniline 362688-26-4,
1-(4-Methylpiperazinyl)-3-aminoisoquinoline
RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

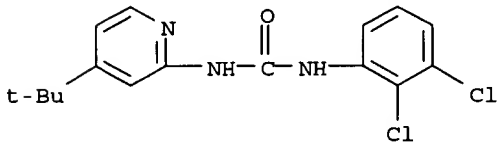
IT 6337-24-2P, 1-Methoxy-4-(4-nitrophenoxy)benzene 13472-85-0P,
5-Bromo-2-methoxypyridine 18994-90-6P, 4-(1-Imidazolylmethyl)-1-nitrobenzene 27692-74-6P, 4-(4-Pyridinylmethyl)aniline 28232-34-0P,
5-Nitro-2-(4-methylphenoxy)pyridine 28232-52-2P, 3-(3-Pyridinyloxy)-1-nitrobenzene 31465-36-8P, 4-(4-Methoxyphenoxy)aniline 32361-76-5P,
3-(4-Nitrobenzyl)pyridine 36089-89-1P, 4-(4-Methylsulfonylphenoxy)-1-nitrobenzene 51834-97-0P, 5-Hydroxy-2-methoxypyridine 62248-47-9P,
4-(4-Butoxyphenyl)thio-1-nitrobenzene 62248-51-5P, 4-(4-Butoxyphenyl)thioaniline 64064-63-7P, 4-(6-Methyl-3-pyridinyloxy)-1-nitrobenzene 70991-08-1P, 4-(2-Pyridinylthio)aniline 92575-23-0P,
3-(4-Pyridinylthio)aniline 116289-71-5P, 3-(3-Pyridinyloxy)aniline
178809-75-1P, 4-[1-Hydroxy-1-(4-pyridyl)methyl]-1-nitrobenzene
220000-87-3P, 2-(N-Methylcarbamoyl)-4-chloropyridine 228401-26-1P,
3-(Trifluoromethyl)-4-(4-pyridinylthio)nitrobenzene 228401-27-2P,
3-(Trifluoromethyl)-4-(4-pyridinylthio)aniline 228401-28-3P,
4-[[4-Phenylthiazol-2-yl]sulfanyl]-1-nitrobenzene 228401-29-4P,

4-[[4-Phenylthiazol-2-yl]sulfanyl]aniline 228401-31-8P, 4-(6-Methyl-3-pyridinyloxy)aniline 228401-32-9P, 4-(3,4-Dimethoxyphenoxy)-1-nitrobenzene 228401-33-0P, 4-(3,4-Dimethoxyphenoxy)aniline 228401-36-3P, 5-Amino-2-(4-methylphenoxy)pyridine Dihydrochloride 228401-37-4P, 4-(3-Thienylthio)-1-nitrobenzene 228401-38-5P, 4-(5-Pyrimidinyloxy)aniline 228401-39-6P, 4-[[2-Methoxypyridin-5-yl]oxy]-1-nitrobenzene 228401-40-9P, 4-(2-Methyl-4-pyridinyloxy)aniline 228401-41-0P, Methyl(4-nitrophenyl)-4-pyridylamine 228401-43-2P, 4-(3-Methoxycarbonyl-4-methoxyphenoxy)-1-nitrobenzene 228401-44-3P, 4-(3-Carboxy-4-methoxyphenoxy)-1-nitrobenzene 229003-17-2P, 3-(5-Methyl-3-pyridinyloxy)-1-nitrobenzene 473915-53-6P, 4-(4-tert-Butoxycarbamoylbenzyl)aniline
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

IT **432050-17-4P**, N-(4-tert-Butylpyridyl)-N'-(2,3-dichlorophenyl)urea
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of quinolyl, isoquinolyl or pyridyl-ureas as inhibitors of raf kinase)

RN 432050-17-4 HCAPLUS

CN Urea, N-(2,3-dichlorophenyl)-N'-[4-(1,1-dimethylethyl)-2-pyridinyl]- (9CI) (CA INDEX NAME)



L75 ANSWER 4 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2002:615574 HCAPLUS

DN 137:169425

ED Entered STN: 16 Aug 2002

TI Preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors

IN **Dumas, Jacques**; **Riedl, Bernd**; **Khire, Uday**; **Wood, Jill E.**; Sibley, Robert N.; **Monahan, Mary-Katherine**; Renick, Joel; Gunn, David E.; **Lowinger, Timothy B.**; **Scott, William J.**; **Smith, Roger A.**

PA **Bayer** Corporation, USA

SO PCT Int. Appl., 125 pp.
CODEN: PIXXD2

DT Patent

LA English

IC ICM C07D215-38
ICS C07D401-12; A61K031-4406; A61K031-47; A61P035-00; C07D401-12; C07D215-00; C07D213-00; C07D401-12; C07D215-00; C07D209-00

CC 27-16 (Heterocyclic Compounds (One Hetero Atom))
Section cross-reference(s): 1

FAN.CNT 5

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002062763	A2	20020815	WO 2002-US3361	20020207
	WO 2002062763	A3	20021010		

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PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG,
US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
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BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

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	US 1999-115877P	P	19990113		
	US 1999-257266	B2	19990225		
	US 1999-425228	B2	19991022		
	US 2001-758548	A2	20010112		

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
WO 2002062763	ICM	C07D215-38
	ICS	C07D401-12; A61K031-4406; A61K031-47; A61P035-00; C07D401-12; C07D215-00; C07D213-00; C07D401-12; C07D215-00; C07D209-00
WO 2002062763	ECLA	C07C275/36; C07D295/18B2D; C07D307/14; C07D401/12+215+213; C07D401/12+215+209; C07D401/12+217+213; C07D401/12+213+207; C07C275/40; C07C311/29; C07C317/22; C07D213/75B2; C07D213/82H; C07D215/38C; C07D217/22; C07D295/12A1; C07D295/12B1D4
US 2002165394	NCL	546/143.000; 546/159.000; 546/306.000; 546/014.000
	ECLA	C07C275/28; C07C275/30; C07C275/32; C07C275/36; C07C275/40; C07C311/29; C07C317/22; C07D209/48D5C1; C07D213/75B2; C07D213/75D3; C07D213/81E; C07D213/82H; C07D215/38C; C07D295/12A1; C07D295/12B1D4; C07D295/18B2D; C07D307/14; C07D401/12+215+213; C07D401/12+215+209; C07D401/12+217+213; C07D401/12+213+207

OS MARPAT 137:169425

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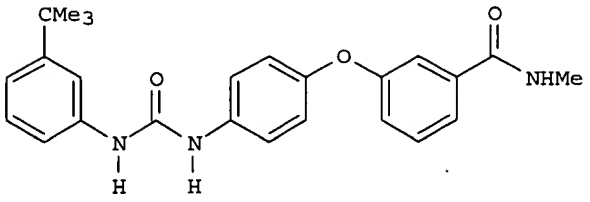

II

AB Title compds., e.g., RNHCONHZOR1 [I; R = C6H4(CMe3)-3, 2-methoxy-5-trifluoromethylphenyl, 4-chloro-3-trifluoromethylphenyl, 2-methoxy-3-quinolyl, etc.; R1 = (un)substituted acylphenyl, -acylpyridinyl, etc.; Z = (un)substituted 1,3- or -1,4-phenylene] were prepared Thus, 4-(H2N)C6H4OC6H4(CONHMe)-4 (preparation given) was condensed with 3-(Me3C)C6H4NH2 and CO(OCCl3)2 to give title compound II. Data for biol. activity of title compds. were given.

ST acylphenoxyphenylurea prepn raf kinase inhibitor; antitumor agent acylphenoxyphenylurea prepn

IT Antitumor agents
Human
(preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors)

IT Neoplasm
(treatment; preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors)

IT 139691-76-2, Raf kinase
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(mediated disorders; treatment; preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors)

IT 228418-48-2P 284461-33-2P 284461-34-3P 284461-35-4P 284461-36-5P
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447457-08-1P 447457-09-2P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors)

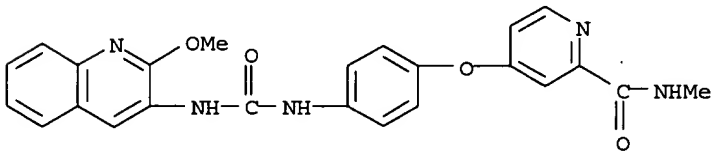
IT 75-31-0, Isopropylamine, reactions 98-58-8, 4-Bromobenzenesulfonyl chloride 98-98-6, Picolinic acid 99-93-4, p-Hydroxyacetophenone 99-98-9, 4-(Dimethylamino)aniline 100-51-6, Benzyl alcohol, reactions 101-79-1, 4-(4-Chlorophenoxy)aniline 106-50-3, p-Phenylenediamine, reactions 108-95-2, Phenol, reactions 109-85-3, 2-Methoxyethylamine 123-30-8, 4-Aminophenol 123-54-6, Acetylacetone, reactions 320-51-4, 4-Chloro-3-(trifluoromethyl)aniline 327-78-6, 4-Chloro-3-(trifluoromethyl)phenyl isocyanate 349-65-5, 2-Methoxy-5-(trifluoromethyl)aniline 350-46-9, 1-Fluoro-4-nitrobenzene 371-40-4, 4-Fluoroaniline 393-36-2, 4-Bromo-3-(trifluoromethyl)aniline 462-08-8, 3-Aminopyridine 490-79-9, 2,5-Dihydroxybenzoic acid 591-27-5, 3-Aminophenol 610-35-5, 4-Hydroxyphthalic acid 619-08-9, 2-Chloro-4-nitrophenol 626-61-9, 4-Chloropyridine 883-99-8, Methyl 3-hydroxy-2-naphthoate 1121-78-4, 5-Hydroxy-2-methylpyridine 1193-02-8, 4-Aminothiophenol 1664-40-0, N-Phenylethylenediamine 1877-71-0, Mono-Methyl isophthalate 2038-03-1, 4-(2-Aminoethyl)morpholine 2524-67-6, 4-Morpholinoaniline 2835-95-2, 5-Amino-2-methylphenol 2835-99-6, 4-Amino-3-methylphenol 2905-24-0, 3-Bromobenzenesulfonyl chloride 3535-88-4, 5-tert-Butyl-2-methoxyaniline 3964-52-1, 4-Amino-2-chlorophenol 4548-45-2, 2-Chloro-5-nitropyridine 4795-29-3, Tetrahydrofurfurylamine 5369-19-7, 3-tert-Butylaniline 6310-19-6, 2-Nitro-4-tert-butylaniline 6628-77-9, 5-Amino-2-methoxypyridine 6927-86-2, 4-(4-Acetylphenoxy)aniline hydrochloride 7781-98-8, Ethyl 3-hydroxybenzoate 13154-24-0, Triisopropylsilyl chloride 22948-02-3, 3-Aminothiophenol 25900-61-2, 3-(Methylcarbamoyl)aniline 26116-12-1, 2-(Aminomethyl)-1-ethylpyrrolidine 27578-60-5, 1-(2-Aminoethyl)piperidine 30766-22-4, Methyl 5-hydroxynicotinate 30806-83-8, Ethyl 4-isocyanatobenzoate 33252-26-5, 2-Amino-4-tert-butylpyridine 34803-66-2, N-(2-Pyridyl)piperazine 36265-31-3, 4-(4-(Methylthio)phenoxy)-1-nitrobenzene 106164-64-1 150009-83-9, 3-Amino-2-methoxyquinoline 284462-71-1 284462-72-2, 3-Chloro-6-(N-acetylamino)-4-(trifluoromethyl)anisole 284462-73-3, 4-Chloro-N-(2-hydroxyethyl)pyridine-2-carboxamide 447457-10-5
RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors)

IT 883-62-5P, 3-Methoxy-2-naphthoic acid 13041-60-6P, Methyl 3-methoxy-2-naphthoate 27237-21-4P, 4-(3-Carboxyphenoxy)-1-nitrobenzene 36089-89-1P, 4-(4-(Methylsulfonyl)phenoxy)-1-nitrobenzene 41513-02-4P, 4-Bromo-3-(trifluoromethyl)phenyl isocyanate 50727-06-5P, 5-Hydroxyisoindoline-1,3-dione 51727-15-2P, 4-Chloropyridine-2-carbonyl chloride hydrochloride 54579-63-4P, 4-(3-Carboxyphenoxy)aniline 64064-63-7P, 4-((2-Methylpyridin-5-yl)oxy)-1-nitrobenzene 67291-63-8P, 2-Amino-3-methoxynaphthalene 71708-64-0P 77992-50-8P, 4-Bromo-3-(trifluoromethyl)aniline hydrochloride 119431-22-0P, 3-Chloro-4-(2,2,2-trifluoroacetylamino)phenol 153435-79-1P, N-Methyl-3-bromobenzenesulfonamide 176977-85-8P, Methyl 4-chloropyridine-2-carboxylate hydrochloride 220000-87-3P 228401-15-8P 228401-43-2P, 4-(3-Methoxycarbonyl-4-methoxyphenoxy)-1-nitrobenzene 228401-44-3P, 4-(3-Carboxy-4-methoxyphenoxy)-1-nitrobenzene 252061-66-8P, 5-Hydroxyisoindolin-1-one 284462-37-9P 284462-38-0P, 5-(4-Nitrophenoxy)isoindoline-1,3-dione 284462-39-1P, 5-(4-Aminophenoxy)isoindoline-1,3-dione 284462-40-4P, 1-(4-tert-Butyl-2-nitrophenyl)-2,5-dimethylpyrrole 284462-41-5P 284462-42-6P 284462-43-7P 284462-44-8P 284462-45-9P, 4-Chloro-2-methoxy-5-(trifluoromethyl)aniline 284462-46-0P, 4-[3-(N-Methylcarbamoyl)-4-methoxyphenoxy]-1-nitrobenzene 284462-47-1P, 4-[3-(N-Methylcarbamoyl)-4-methoxyphenoxy]aniline 284462-48-2P, 5-(4-Nitrophenoxy)-2-methylisoindoline-1,3-dione 284462-49-3P, 5-(4-Aminophenoxy)-2-methylisoindoline-1,3-dione 284462-51-7P, 4-Chloro-2-[N-(2-morpholin-4-ylethyl)carbamoyl]pyridine 284462-52-8P 284462-53-9P, 4-(1-Oxoisoindolin-5-yloxy)-1-nitrobenzene 284462-54-0P, 4-(1-Oxoisoindolin-5-yloxy)aniline 284462-55-1P, 4-(3-Ethoxycarbonylphenoxy)-1-nitrobenzene 284462-56-2P 284462-57-3P 284462-58-4P 284462-59-5P 284462-60-8P, 4-[3-(N-Methylsulfamoyl)phenoxy]-1-nitrobenzene 284462-61-9P, 4-[3-(N-Methylsulfamoyl)phenoxy]aniline 284462-62-0P 284462-63-1P, 4-Chloro-N-[2-(triisopropylsilyloxy)ethyl]pyridine-2-carboxamide 284462-64-2P 284462-65-3P, 4-((2-Methoxycarbonylpyridin-5-yl)oxy)-1-nitrobenzene 284462-66-4P 284462-67-5P 284462-68-6P 284462-69-7P 284462-97-1P 432050-20-9P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors)

IT **432050-22-1P**
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of N-aryl-N'-[(acylphenoxy)phenyl]ureas as raf kinase inhibitors)

RN 432050-22-1 HCAPLUS

CN 2-Pyridinecarboxamide, 4-[4-[[[(2-methoxy-3-quinolinyl)amino]carbonyl]amino]phenoxy]-N-methyl- (9CI) (CA INDEX NAME)



L75 ANSWER 5 OF 5 HCAPLUS COPYRIGHT 2005 ACS on STN

AN 2002:409267 HCAPLUS

DN 137:6098

ED Entered STN: 31 May 2002

TI Heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors

IN **Dumas, Jacques; Riedl, Bernd; Khire, Uday;** Sibley, Robert N.; **Hatoum-Mokdad, Holia; Monahan,**

Mary-katherine; Gunn, David E.; Lowinger, Timotthy B.; Scott, William J.; Smith, Roger A.; Wood, Jill E.
PA Bayer Corporation, USA
SO U.S. Pat. Appl. Publ., 39 pp., Cont.-in-part of U. S. Ser. No. 778,039.
CODEN: USXXCO
DT Patent
LA English
IC ICM A61K031-506
ICS A61K031-501; A61K031-497; A61K031-4725; A61K031-4709
INCL 514310000
CC 27-17 (Heterocyclic Compounds (One Hetero Atom))
Section cross-reference(s): 1
FAN.CNT 5

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PI	US 2002065296	A1	20020530	US 2001-838286	20010420
	US 2003139605	A1	20030724	US 2002-71248	20020211
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	US 1999-257265	B1	19990225		
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	US 1999-257266	B2	19990225		
	US 1999-425228	B1	19991022		
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	US 2001-948915	A1	20010910		
	WO 2002-US12064	W	20020417		

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
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US 2002065296	NCL	514/310.000; 514/313.000; 514/336.000; 514/337.000; 514/252.030; 514/252.040; 514/255.050; 514/256.000
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US 2003139605	NCL	546/291.000
	ECLA	A61K031/17; C07C311/29; C07C317/22; C07D209/48D5C1; C07D213/75D3; C07D213/81E; C07D295/12A1; C07D295/12B1D4; C07D295/18B2D; A61K031/18; A61K031/24; A61K031/341; A61K031/40+A; A61K031/4035; A61K031/44+A; A61K031/4439; A61K031/4453; A61K031/495+A; A61K031/496; A61K031/5375; A61K031/5377; C07C275/28; C07C275/30; C07C275/32; C07C275/36; C07C275/40
WO 2002085859	ECLA	A61K031/44; A61K031/47; A61K031/4709; A61K031/4725;

C07D213/75D3; C07D215/38C; C07D217/22;
C07D401/12+215+213; C07D401/12+217+213
JP 2004537511 FTERM 4C031/JA09; 4C034/AL05; 4C055/AA01; 4C055/BA01;
4C055/BA02; 4C055/BA53; 4C055/BB17; 4C055/CA01;
4C055/DA06; 4C055/DA28; 4C055/DA42; 4C055/DA47;
4C055/DB10; 4C055/DB17; 4C055/EA01; 4C063/AA01;
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4C086/MA01; 4C086/MA04; 4C086/NA14; 4C086/ZA01;
4C086/ZA02; 4C086/ZA36; 4C086/ZA45; 4C086/ZA54;
4C086/ZA59; 4C086/ZA67; 4C086/ZA68; 4C086/ZA75;
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4C086/ZA97; 4C086/ZB02; 4C086/ZB05; 4C086/ZB11;
4C086/ZB13; 4C086/ZB15; 4C086/ZB26; 4C086/ZB33;
4C086/ZB35; 4C086/ZC21; 4C086/ZC35

OS MARPAT 137:6098

AB This invention relates to the use of a group of heteroaryl ureas (I; for example, N-(2-methoxy-3-quinolyl)-N'-[4-[3-(N-methylcarbamoyl)phenoxy]phenyl]urea) containing N in treating p38 mediated diseases, and pharmaceutical compns. for use in such therapy. I is A-NHC(O)NH-B or a pharmaceutically acceptable salt thereof, wherein A is a substituted or unsubstituted pyridyl, quinolinyl or isoquinolinyl group, B is a substituted or unsubstituted, up to tricyclic aryl or heteroaryl moiety of up to 50 C atoms with a cyclic structure bound directly to N, containing at least 5 cyclic members with 0-4 members of groups consisting of N, O and S. Information about the substituents for A and B are given in the claims. Although the methods of preparation are not claimed, 37 example prepns. are included as well as examples of preparation of intermediates. No pharmacol. data is included.

ST nitrogen heteroaryl urea prepn p38 kinase inhibitor; pyridyl urea prepn p38 kinase inhibitor; quinolyl urea prepn p38 kinase inhibitor; isoquinolyl urea prepn p38 kinase inhibitor

IT Infection
(Chagas' disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Inflammation
(Crohn's disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Intestine, disease
(Crohn's; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Disease, animal
(Jarisch-Herxheimer reaction; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Malaria
(Plasmodium falciparum malaria and cerebral malaria; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treating)

IT Antimalarials
(Plasmodium falciparum malaria and cerebral malaria; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Toxins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(Shiga-like toxin, effects of toxins from Escherichia coli infection; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Respiratory distress syndrome
(adult; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Hepatitis
(alc.; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Transplant rejection

(allotransplant; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Lung
(alveolus, injury; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Antiarteriosclerotics
(antiatherosclerotics; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)
IT Aneurysm
(aortic; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Meningitis
(bacterial; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Necrosis
(bowel; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Bronchi, disease
Inflammation
(bronchitis, obliterative; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Injury
(cerebral; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Pneumoconiosis
(coal worker's; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Eye, disease
(cornea, ulcer; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Ulcer
(corneal; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Radiation
(damage, injury/toxicity following administration of monoclonal antibodies; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Cartilage, disease
(degeneration; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Multiple sclerosis
(demyelation and oligiodendrocyte loss in; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Liver, disease
(during acute inflammation; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Toxins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(enterotoxin A, effects of toxins from Staphylococcus infection; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Skin, disease
(epidermolysis bullosa, dystrophobic; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Liver, disease
(failure; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Lung, disease
(fibrosis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Nervous system agents
(for demyelating disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)
IT Wound healing
(impaired wound healing in infection; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Helicobacter pylori
(infection during peptic ulcer disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Borrelia burgdorferi
Cytomegalovirus
Human immunodeficiency virus
Influenza virus
Theiler's murine encephalomyelitis virus
Treponema pallidum
(infections from; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Brain, disease
Reperfusion
(injury; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Leukemia
(lymphocytic, inhibitors; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)
IT Neoplasm
(metastasis, inhibitors; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)
IT Heterocyclic compounds
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(nitrogen, heteroaryl ureas; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)
IT Bone, disease
(osteopenia, mediated by MMP activity; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Inflammation
Pancreas, disease
(pancreatitis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Ulcer
(peptic, Helicobacter pylori infection during; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Osteoporosis
(postmenopausal; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treating)
IT Human
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)
IT Allergy
Alzheimer's disease
Arthritis
Asthma
Diabetes mellitus
Rheumatoid arthritis
Tuberculosis
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treating)
IT Encephalitis
Myelodysplastic syndromes
Periodontium, disease
Psoriasis
Rheumatic fever
Silicosis
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)
IT Allergy inhibitors
Anti-Alzheimer's agents
Anti-infective agents
Anti-inflammatory agents
Antiarthritics

Antiasthmatics
Antibacterial agents
Anticoagulants
Antidiabetic agents
Antirheumatic agents
Antitumor agents
Cardiovascular agents
Contraceptives
Tuberculostatics
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Biliary tract, disease
(primary biliary cirrhosis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Proteins
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(proteinuria; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Fibrosis
Sarcoidosis
(pulmonary; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Injury
(reperfusion; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Bone
(resorption; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Lung, disease
(sarcoidosis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Shock (circulatory collapse)
(septic; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Inflammation
(systemic inflammatory response syndrome; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Lupus erythematosus
(systemic; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Disease, animal
(temporomandibular joint; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Joint, anatomical
(temporomandibular, disease; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Osteoporosis
(therapeutic agents, postmenopausal; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for use as)

IT Shock (circulatory collapse)
(toxic shock syndrome; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Digestive tract, disease
(ulcer, peptic, Helicobacter pylori infection during; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT Inflammation
Intestine, disease
(ulcerative colitis; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors for treatment of)

IT 165245-96-5, p38 Kinase
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(inhibitors; preparation of heteroaryl ureas containing nitrogen hetero-atoms as)

IT 673-09-6P, 4-(4-Pyridylthio)aniline 6337-24-2P, 1-Methoxy-4-(4-nitrophenoxy)benzene 13472-85-0P, 5-Bromo-2-methoxypyridine

18994-90-6P, 4-(1-Imidazolylmethyl)-1-nitrobenzene 27237-21-4P, 4-(3-Carboxyphenoxy)-1-nitrobenzene 27692-74-6P, 4-(4-Pyridinylmethyl)aniline 28232-34-0P, 5-Nitro-2-(4-methylphenoxy)pyridine 28232-52-2P, 3-(3-Pyridinyloxy)-1-nitrobenzene 29264-35-5P, 4-(3-Carboxy-4-hydroxyphenoxy)-1-nitrobenzene 31465-36-8P, 4-(4-Methoxyphenoxy)aniline 32361-76-5P, 3-(4-Nitrobenzyl)pyridine 36089-89-1P, 4-(4-Methylsulfonylphenoxy)-1-nitrobenzene 50727-06-5P, 5-Hydroxyisoindoline-1,3-dione 51727-15-2P, 4-Chloropyridine-2-carbonyl chloride hydrochloride 51834-97-0P, 5-Hydroxy-2-methoxypyridine 56643-85-7P, 4-(1-Imidazolylmethyl)aniline 62248-47-9P, 4-[(4-Butoxyphenyl)thio]-1-nitrobenzene 62248-51-5P, 4-(4-Butoxyphenyl)thioaniline 64064-63-7P, 4-(6-Methyl-3-pyridinyloxy)-1-nitrobenzene 70991-08-1P, 4-(2-Pyridinylthio)aniline 71708-64-0P, 4-[3-(N-Methylcarbamoyl)phenoxy]-1-nitrobenzene 85666-15-5P, 4-[(3-Pyridinyl)methyl]aniline 92575-23-0P, 3-(4-Pyridinylthio)aniline 99586-65-9P, 4-Chloro-2-pyridinecarboxamide 102877-78-1P 116289-71-5P, 3-(3-Pyridinyloxy)aniline 135680-03-4P, 4-(4-tert-Butoxycarbonylaminobenzyl)aniline 176977-85-8P, Methyl 4-chloropyridine-2-carboxylate hydrochloride 178809-75-1P, 4-[Hydroxy(4-pyridyl)methyl]-1-nitrobenzene 220000-87-3P 228401-26-1P, 3-(Trifluoromethyl)-4-(4-pyridinylthio)nitrobenzene 228401-27-2P, 3-(Trifluoromethyl)-4-(4-pyridinylthio)aniline 228401-28-3P, 4-[(4-Phenyl-2-thiazolyl)thio]-1-nitrobenzene 228401-29-4P, 4-[(4-Phenyl-2-thiazolyl)thio]aniline 228401-31-8P, 4-(6-Methyl-3-pyridinyloxy)aniline 228401-32-9P, 4-(3,4-Dimethoxyphenoxy)-1-nitrobenzene 228401-33-0P, 4-(3,4-Dimethoxyphenoxy)aniline 228401-34-1P, 3-(6-Methyl-3-pyridinyloxy)-1-nitrobenzene 228401-35-2P, 3-(6-Methyl-3-pyridinyloxy)aniline 228401-36-3P, 5-Amino-2-(4-methylphenoxy)pyridine Dihydrochloride 228401-37-4P, 4-(3-Thienylthio)-1-nitrobenzene 228401-38-5P, 4-(5-Pyrimidinyloxy)aniline 228401-39-6P, 4-[(2-Methoxy-5-pyridyl)oxy]-1-nitrobenzene 228401-40-9P, 4-(2-Methyl-4-pyridinyloxy)aniline 228401-41-0P, Methyl(4-nitrophenyl)(4-pyridyl)amine 228401-43-2P, 4-(3-Methoxycarbonyl-4-methoxyphenoxy)-1-nitrobenzene 228401-44-3P, 4-(3-Carboxy-4-methoxyphenoxy)-1-nitrobenzene 284462-37-9P, 4-[2-(N-Methylcarbamoyl)-4-pyridyloxy]aniline 284462-38-0P, 5-(4-Nitrophenoxy)isoindoline-1,3-dione 284462-39-1P, 5-(4-Aminophenoxy)isoindoline-1,3-dione 284462-46-0P, 4-[3-(N-Methylcarbamoyl)-4-methoxyphenoxy]-1-nitrobenzene 284462-47-1P, 4-[3-(N-Methylcarbamoyl)-4-methoxyphenoxy]aniline 284462-55-1P, 4-(3-Ethoxycarbonylphenoxy)-1-nitrobenzene 284462-56-2P, 4-(3-N-Methylcarbamoylphenoxy)aniline 284462-78-8P, 3-[2-(N-Methylcarbamoyl)-4-pyridyloxy]aniline 284462-79-9P, 3-(2-Carbamoyl-4-pyridyloxy)aniline 284462-80-2P, 4-(2-Carbamoyl-4-pyridyloxy)aniline 284462-84-6P, 4-(4-Methylsulfonylphenoxy)aniline 432050-13-0P, 4-(3-Thienylthio)aniline 432050-14-1P, 4-[(2-Methoxy-5-pyridyl)oxy]aniline 432050-15-2P, Methyl(4-aminophenyl)(4-pyridyl)amine 432050-16-3P, 4-[Hydroxy(4-pyridyl)methyl]aniline
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(intermediate; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

IT 284461-54-7P, N-[2-Methoxy-5-(trifluoromethyl)phenyl]-N'-[4-(1,3-dioxoisoindolin-5-yloxy)phenyl]urea 284670-98-0P, N,N'-Bis[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea **432050-17-4P** 432050-18-5P 432050-19-6P, N,N'-Bis(2-methoxy-3-quinolinyl)urea 432050-20-9P 432050-21-0P, N-[5-Trifluoromethyl-2-pyridyl]-N'-[3-(4-pyridylthio)phenyl]urea **432050-22-1P**, N-(2-Methoxy-3-quinolinyl)-N'-[4-(2-(N-Methylcarbamyl)-4-pyridyloxy)phenyl]urea **432050-23-2P**, N-(2-Methoxy-3-quinolyl)-N'-[4-[3-(N-methylcarbamoyl)phenoxy]phenyl]urea **432050-24-3P**, N-(2-Methoxy-3-quinolyl)-N'-[4-(2-carbamoyl-4-pyridyloxy)phenyl]urea **432050-25-4P**, N-(2-Methoxy-3-quinolyl)-N'-[3-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea **432050-26-5P**, N-(2-Methoxy-3-quinolyl)-N'-[3-(2-carbamoyl-4-pyridyloxy)phenyl]urea **432050-27-6P**, N-(2-Methoxy-3-quinolyl)-N'-[4-[3-(N-isopropylcarbamoyl)phenoxy]phenyl]urea 432050-28-7P,

N-(2-Methoxy-3-quinolyl)-N'-[4-[4-methoxy-3-(N-methylcarbamoyl)phenoxy]phenyl]urea **432050-29-8P**, N-(3-Isoquinolyl)-N'-[4-[2-(N-methylcarbamoyl)-4-pyridyloxy]phenyl]urea **432050-30-1P**, N-(4-tert-Butyl-2-pyridinyl)-N'-(4-methylphenyl)urea **432050-31-2P**, N-(4-tert-Butyl-2-pyridinyl)-N'-(4-fluorophenyl)urea **432050-32-3P**, N-(4-tert-Butyl-2-pyridinyl)-N'-(1-naphthyl)urea **432050-33-4P**, N-(4-tert-Butyl-2-pyridinyl)-N'-[4-(4-methoxyphenoxy)phenyl]urea 432050-34-5P, N-(5-Trifluoromethyl-2-pyridinyl)-N'-[4-(4-pyridylmethyl)phenyl]urea **432050-35-6P**, N-(3-Isoquinolyl)-N'-(4-methylphenyl)urea **432050-36-7P**, N-(3-Isoquinolyl)-N'-(4-fluorophenyl)urea **432050-37-8P**, N-(3-Isoquinolyl)-N'-(2,3-dichlorophenyl)urea **432050-38-9P**, N-(3-Isoquinolyl)-N'-(1-naphthyl)urea **432050-39-0P**, N-(3-Isoquinolyl)-N'-[4-(4-pyridinylmethyl)phenyl]urea **432050-40-3P**, N-(3-Quinolyl)-N'-[4-(4-pyridinylmethyl)phenyl]urea 432050-41-4P, N-(4-tert-Butyl-2-pyridyl)-N'-(4-(4-methylphenoxy)phenyl)urea 432050-42-5P, N-(4-tert-Butyl-2-pyridyl)-N'-(4-(4-pyridyloxy)phenyl)urea 432050-43-6P, N-(4-tert-Butyl-2-pyridyl)-N'-(4-(4-pyridinylthio)phenyl)urea 432050-44-7P, N-(4-tert-Butyl-2-pyridyl)-N'-(3-(4-pyridinylthio)phenyl)urea 432050-45-8P 432050-46-9P 432050-47-0P 432050-48-1P 432050-49-2P 432050-50-5P 432050-51-6P, N-(1-(4-Methyl-1-piperazinyl)isoquinol-3-yl)-N'-(4-((4-pyridyl)methyl)phenyl)urea 432050-52-7P, N-(Isoquinol-3-yl)-N'-(4-(3-(methylcarbamoyl)phenoxy)phenyl)urea 432050-53-8P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

IT 75-31-0, Isopropylamine, reactions 86-84-0, 1-Naphthyl isocyanate 98-98-6, Picolinic acid 100-11-8, 4-Nitrobenzyl bromide 100-15-2, N-Methyl-4-nitroaniline 101-77-9, 4,4'-Methylenedianiline 101-79-1, 4-(4-Chlorophenoxy)aniline 106-44-5, 4-Methylphenol, reactions 109-00-2, 3-Hydroxypyridine 123-30-8, 4-Aminophenol 139-59-3, 4-Phenoxyaniline 150-76-5, 4-Methoxyphenol 288-32-4, Imidazole, reactions 350-46-9, 1-Fluoro-4-nitrobenzene 400-74-8, 2-Fluoro-5-nitrobenzotrifluoride 580-17-6, 3-Aminoquinoline 585-79-5, 1-Bromo-3-nitrobenzene 591-27-5, 3-Aminophenol 610-35-5, 4-Hydroxyphthalic acid 620-95-1, 3-Benzylpyridine 622-58-2, 4-Tolyl isocyanate 624-28-2, 2,5-Dibromopyridine 626-61-9, 4-Chloropyridine 626-64-2, 4-Hydroxypyridine 872-31-1, 3-Bromothiophene 1083-48-3, 4-(4-Nitrobenzyl)pyridine 1121-78-4, 5-Hydroxy-2-methylpyridine 1193-02-8, 4-Aminothiophenol 1195-45-5, 4-Fluorophenyl isocyanate 1849-36-1, 4-Nitrothiophenol 2033-89-8, 3,4-Dimethoxyphenol 2103-88-0, 2-Mercapto-4-phenylthiazole 3678-63-5, 4-Chloro-2-methylpyridine 4548-45-2, 2-Chloro-5-nitropyridine 4556-23-4, 4-Mercaptopyridine 4595-59-9, 5-Bromopyrimidine 7379-35-3, 4-Chloropyridine hydrochloride 7781-98-8, Ethyl 3-hydroxybenzoate 16588-75-3, 2-Methoxy-5-(trifluoromethyl)phenyl isocyanate 21101-60-0, 4-(4-Nitrophenylthio)phenol 22948-02-3, 3-Aminothiophenol 24424-99-5, Di-tert-butyl dicarbonate 25267-27-0, Iodobutane 25475-67-6, 3-Aminoisoquinoline 27163-00-4, 4-[(4-Methoxyphenyl)methylamino]aniline 33252-26-5, 2-Amino-4-tert-butylpyridine 36265-31-3, 4-(4-Methylthiophenoxy)-1-nitrobenzene 41195-90-8, 2,3-Dichlorophenyl isocyanate 41295-20-9, 4-(4-Methylphenoxy)aniline 53750-66-6, 4-Chloropyridine-2-carbonyl chloride 73322-01-7, 4-(2-Pyridinylthio)-1-nitrobenzene 74784-70-6, 2-Amino-5-(trifluoromethyl)pyridine 150009-83-9, 3-Amino-2-methoxyquinoline 170893-64-8, 4-(4-Pyridylcarbonyl)aniline 362688-26-4, 1-(4-Methylpiperazinyl)-3-aminoisoquinoline

RL: RCT (Reactant); RACT (Reactant or reagent)

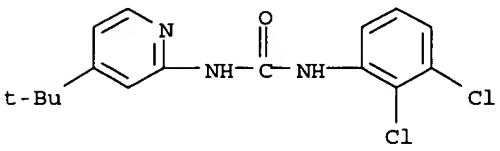
(reactant; preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

IT **432050-17-4P**

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(Uses)
(preparation of heteroaryl ureas containing nitrogen hetero-atoms as p38 kinase inhibitors)

RN 432050-17-4 HCAPLUS
CN Urea, N-(2,3-dichlorophenyl)-N'-[4-(1,1-dimethylethyl)-2-pyridinyl]- (9CI) (CA INDEX NAME)



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L76 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2005 ACS on STN
AN 2004:701815 HCAPLUS
DN 141:185104
ED Entered STN: 27 Aug 2004
TI Compositions, combinations, and methods for treating cardiovascular conditions and other associated conditions
IN Rudolph, Amy E.; Rocha, Ricardo; Carretero, Oscar
PA USA
SO U.S. Pat. Appl. Publ., 107 pp.
CODEN: USXXCO
DT Patent
LA English
IC ICM A61K031-415
ICS A61K031-401
INCL 514406000; 514423000
CC 1-8 (Pharmacology)
FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI US 2004167197	A1	20040826	US 2004-788220	20040226
WO 2004075852	A2	20040910	WO 2004-US5609	20040226

W: AE, AE, AG, AL, AL, AM, AM, AM, AT, AT, AU, AZ, AZ, BA, BB, BG, BG, BR, BR, BW, BY, BY, BZ, BZ, CA, CH, CN, CN, CO, CO, CR, CR, CU, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EC, EC, EE, EE, EG, ES, ES, FI, FI, GB, GD, GE, GE, GH, GM, HR, HR, HU, HU, ID, IL, IN, IS, JP, JP, KE, KE, KG, KG, KP, KP, KP, KR, KR, KZ, KZ, KZ, LC, LK, LR, LS, LS, LT, LU, LV, MA, MD, MD, MG, MK, MN, MW, MX, MX, MZ, MZ, NA, NI
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

WO 2004075857	A2	20040910	WO 2004-US5799	20040226

W: AE, AE, AG, AL, AL, AM, AM, AM, AT, AT, AU, AZ, AZ, BA, BB, BG, BG, BR, BR, BW, BY, BY, BZ, BZ, CA, CH, CN, CN, CO, CO, CR, CR, CU, CU, CZ, CZ, DE, DE, DK, DK, DM, DZ, EC, EC, EE, EE, EG, ES, ES, FI, FI, GB, GD, GE, GE, GH, GM, HR, HR, HU, HU, ID, IL, IN, IS, JP, JP, KE, KE, KG, KG, KP, KP, KP, KR, KR, KZ, KZ, KZ, LC, LK, LR, LS, LS, LT, LU, LV, MA, MD, MD, MG, MK, MN, MW, MX, MX, MZ, MZ, NA, NI
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRAI US 2003-450529P P 20030226

CLASS

PATENT NO.	CLASS	PATENT FAMILY CLASSIFICATION CODES
US 2004167197	ICM	A61K031-415
	ICS	A61K031-401
	INCL	514406000; 514423000
US 2004167197	NCL	514/406.000; 514/423.000

AB This invention is directed generally to a method for treating a pathol. condition (particularly a cardiovascular condition (e.g., hypertension or heart failure) or a condition associated with a cardiovascular condition) using a p38-kinase inhibitor (e.g., a p38-kinase-inhibiting substituted pyrazole), and specifically a combination comprising a p38-kinase inhibitor with an angiotensin-converting-enzyme inhibitor (or "ACE inhibitor") for treating a cardiovascular condition. This invention also is directed generally to combinations comprising a p38-kinase inhibitor, and specifically to combinations comprising a p38-kinase inhibitor with an angiotensin-converting-enzyme inhibitor. This invention is further directed generally to pharmaceutical compns. comprising a p38-kinase inhibitor, and more specifically to compns. comprising the above-described combinations.

ST cardiovascular hypertension heart failure p38 kinase pyrazole ACE inhibitor

IT Heart, disease
(arrhythmia; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Fibrosis
Ischemia
(cardiac; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Heart, disease
(cardiomyopathy; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Brain, disease
(cerebrovascular; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Anti-ischemic agents
Antiarrhythmics
Antihypertensives
Blood vessel, disease
Cardiovascular agents
Cardiovascular system, disease
Combination chemotherapy
Drug interactions
Edema
Hypertension
Kidney, disease
(compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Surgery
(coronary angioplasty; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Thrombosis
(coronary arterial; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Artery
(coronary, angioplasty; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Artery, disease
(coronary, thrombosis; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Artery
(coronary; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT Blood vessel, disease
(endothelium; compns., combinations, and methods for treating

cardiovascular conditions and other associated conditions)
IT Heart, disease
(failure; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Heart, disease
(fibrosis; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Heart, disease
(infarction; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Heart, disease
(ischemia; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Heart, disease
Inflammation
(myocarditis; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Nerve, disease
(neuropathy; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Eye, disease
(retinopathy; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Endothelium
(vascular, disease; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Blood vessel, disease
Inflammation
(vasculitis; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Heart, disease
(ventricle, hypertrophy; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT Hypertrophy
(ventricular; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT 165245-96-5, p38-Kinase
RL: BSU (Biological study, unclassified); BIOL (Biological study)
(compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)
IT 39698-78-7, Saralasin acetate 62571-86-2, Captopril 72873-74-6
74258-86-9, Alacepril 75847-73-3, Enalapril 76420-72-9, Enalaprilat
76547-98-3, Lisinopril 82768-85-2, Quinaprilat 82834-16-0, Perindopril
83435-66-9, Delapril 83647-97-6, Spirapril 85441-61-8, Quinapril
85856-54-8, Moveltipril 86541-75-5, Benazepril 87333-19-5, Ramipril
87679-37-6, Trandolapril 88768-40-5, Cilazapril 89371-37-9, Imidapril
95399-71-6, Fosinoprilat 98048-97-6, Fosinopril 103775-10-6, Moexipril
111223-26-8, Cero-napril 111902-57-9, Temocapril 152121-30-7
152121-47-6 164301-51-3 165806-09-7 165806-34-8 165806-48-4
165806-51-9 165806-53-1 180869-32-3 188352-45-6 189442-43-1
193746-75-7 200801-85-0 209410-46-8 215303-72-3 215306-39-1
218145-98-3 218160-26-0 218162-38-0 219138-27-9 249937-52-8
250123-27-4 271576-76-2 271576-77-3 271576-80-8 271576-82-0
271576-83-1 271576-87-5 271576-89-7 271576-90-0 271576-91-1
271576-92-2 271576-93-3 271577-04-9 271577-08-3 285983-48-4
290357-24-3 304439-93-8 321351-00-2 335652-44-3 431062-04-3
432042-02-9 **432050-23-2** 635724-76-4 635724-78-6
635724-79-7 635724-80-0 635724-81-1 635724-82-2 635724-89-9
635724-91-3 635724-92-4 635724-94-6 635724-95-7 635724-96-8
635724-98-0 635724-99-1 635725-04-1 635725-05-2 635725-07-4
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635725-44-9 635725-45-0 635725-46-1 635725-47-2 635725-48-3
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635725-65-4 635725-66-5 635725-67-6 635725-68-7 635725-69-8
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635725-85-8 635725-86-9 635725-87-0 635725-88-1 635725-89-2
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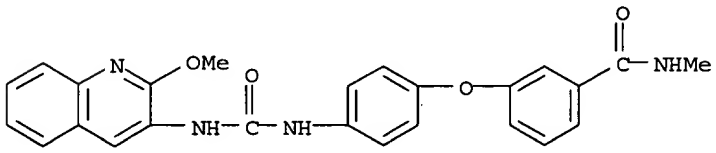
740845-71-0 740845-72-1 740845-73-2 740845-74-3 740845-75-4
740845-76-5 740845-77-6 740845-78-7 740845-79-8 740845-80-1
740845-81-2
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT 9015-82-1, Angiotensin-converting enzyme
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(inhibitors; compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

IT **432050-23-2**
RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(compns., combinations, and methods for treating cardiovascular conditions and other associated conditions)

RN 432050-23-2 HCAPLUS
CN Benzamide, 3-[4-[[[(2-methoxy-3-quinolinyl)amino]carbonyl]amino]phenoxy]-N-methyl- (9CI) (CA INDEX NAME)



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